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Isolation of Fungi From Inoculated Guinea Pigs
Expert Committee on Tuberculosis
Incidence of Communicable Diseases in the U. S.



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^{*}This is the twenty-seventh of a series of special issues of Public Health Reports devoted exclusively to tuberculosis control, which will appear the first week of each month. The series began with the Mar. 1, 1946 issue. The articles in these special issues are reprinted as extracts from the Public Health Reports. Effective with the July 5, 1946, issue, these extracts may be purchased from the Superintendent of Documents, Government Printing Office, Washington 25, D. C., for 10 cents a single copy. Subscriptions are obtainable at \$1.00 per year; \$1.25 foreign.

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FURTHER STUDY OF BCG VACCINATION

At the 1946 conference on BCG vaccination, held in Washington in September of that year, certain recommendations were made which were reported in the March 7, 1947, issue of Public Health Reports. Since that time, nation-wide interest in this immunizing procedure has grown to an unprecedented level of intensity. So great, indeed, had been the demand from every quarter for additional information concerning the course of BCG investigations that an evaluation and review of recent work in the field had become increasingly appropriate. To that end, a second conference convened in New York City on March 9, 1948, attended by the same group which had met in 1946 to formulate the original policy.

In considering the present status of BCG, the New York conference recognized that many fundamental problems are yet to be resolved concerning the vaccination technique and the vaccine itself. Accordingly, the committee of specialists recommended that the entire problem of BCG continue to receive further attention in the form of intensified research and critical study, and that its application in this country be restricted to certain groups and circumstances.

The decision to limit the use of BCG for the present would appear to be wholly sound and practical. In certain other diseases, such as smallpox, where the desirability of vaccination procedures has been so clearly established, it would be difficult, indeed, to conceive of control without the practice of immunization. In tuberculosis, however, where our fund of knowledge regarding immunization is as yet incomplete, vaccination must continue to be carried out under appropriate safeguards.

The results of careful study leave little doubt that BCG is harmless. As for the degree of immunity which the agent confers, however, there has been very little unanimity of opinion. Although it has become increasingly evident that properly administered BCG vaccination does increase individual resistance to tuberculous infection, there is still a singular lack of definitive information concerning the exact duration of the protection bestowed by the agent. The reason for this is simply that to date, there has been no reliable study of sufficient duration to permit precise evaluation of the vaccine's effectiveness over long periods of time.

Should further study demonstrate that BCG vaccination confers only short-term protection, this would not necessarily argue against its use. In such an eventuality, revaccination might serve to reestablish temporary protection, as is sometimes necessary in the case of other immunizing agents when the immunity conferred by such agents wanes with the passage of time. It can be reasonably expected that further research will provide solutions to these and other prob-

lems now confronting BCG investigators.

Regardless of the ultimate place of this immunizing agent in the nation's tuberculosis control program, however, it is extremely unlikely that BCG will ever obviate entirely the necessity for the conscientious prosecution of proved control measures. These must be exploited to the fullest in order to secure past gains and to meet extant needs. Indeed, there is a considerable body of opinion in this country which holds that modern methods of diagnosis and treatment make possible the effective control of the disease without resort to vaccination procedures, especially in areas of low prevalence which possess adequate diagnostic and medical-care facilities. In such locales, the widespread use of BCG could be expected to make relatively little contribution to the control of the disease at this time. In some respects, indeed, it might serve only to confuse existing epidemiological patterns, particularly with reference to the alteration of tuberculin sensitivity. From this point of view, one can appreciate the concern expressed in some quarters that the present balance of tuberculosis control may be disturbed or possibly dislocated by the unqualified acceptance of an insufficiently tried alternative to present control measures.

Therefore, until all the problems of BCG vaccination as a tuberculosis control technique are resolved, and until we can be certain of the implications of a change in emphasis at this time, we have no right to relax our efforts in the application of measures which long experience has shown to be eminently practical and profitable. To do so prematurely might conceivably prove wasteful, if not disastrous, for when incompletely proved techniques are hastily adopted, it frequently happens that "those who cannot remember the past are condemned to repeat it." True progress lies not in that direction.

Francis J. Weber, Medical Director, Chief, Tuberculosis Control Division. ta

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STUDIES OF FUNGUS ANTIGENS. III.

Sensitization of Normal Animals With Skin Test Antigens 1

By ARDEN HOWELL, Jr., Ph. D., Senior Mycologist, Public Health Service

INTRODUCTION

It has been shown (1, 2) that repeated intradermal injections of 0.1 ml. of 1–10 and 1–100 dilutions of each of several lots of histoplasmin and blastomycin into normal guinea pigs does not sensitize these animals. It has also been shown that coccidioidin (3) and other skin test antigens (4, 5, 6), injected subcutaneously or intraperitoneally, may sensitize guinea pigs. The following experiment was undertaken, therefore, to determine the effects of repeated intradermal injections of more concentrated dilutions of histoplasmin, blastomycin, and other fungus antigens on normal guinea pigs.

MATERIALS AND METHODS

Each of 20 normal albino guinea pigs was tested simultaneously with a skin test dose (0.1 ml.) of undiluted histoplasmin, a 1–10, and a 1–100 dilution of histoplasmin, lots H–15 and H–6 (2). At the same time, 10 of these were tested with similar doses of a 1–100 dilution of a heat-killed yeast-phase antigen of *Histoplasma capsulatum* (2); the other 10 with a 1–10 dilution of this antigen. After 35 days, 19 of these animals were retested with the same amount of the same 3 dilutions of histoplasmin, lot H–15, and with a 1–100 dilution of the yeast-phase antigen. At intervals thereafter, the animals were retested with a 1–100 dilution of lot H–15 histoplasmin.

A second group of 10 normal albino guinea pigs were tested simultaneously with 0.1 ml. of undiluted, 1–10, and 1–100 dilutions of blastomycin, lots B–2 and B–7, and a 1–100 dilution of a heat-killed yeast-phase antigen of *Blastomyces dermatitidis* (2). Thirty-five days later the tests were repeated with the yeast-phase antigen and lot B–7. At intervals thereafter, the animals were retested with a 1–100 dilution of lot B–7 blastomycin.

A third group, composed of 38 normal albino guinea pigs, were tested with a skin test dose of each of several dilutions of an autoclaved filtrate antigen prepared from *Candida* (*Monilia*) albicans (7). Fifteen of these were tested with a 1–100 dilution of this antigen, 11 with a 1–10 and a 1–100 dilution, and 12 with undiluted, a 1–10, and a 1–100 dilution. Fifteen days later these tests were repeated on the same animals in the same manner.

A fourth group, composed of 20 normal albino guinea pigs were skin-tested with 0.1 ml. of a 1-100 dilution of a heat-killed suspension of *Candida* (*Monilia*) albicans ("vaccine-type" antigen) and a 1-100 dilution of histoplasmin, lot H-15. Fifteen days later these tests

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From the Office of Field Studies, Tuberculosis Control Division.

were repeated. After 182 days, 19 of these 20 were retested with 0.1 ml. of the autoclaved filtrate antigen of *C. albicans* and lot H-15 histoplasmin. Sixteen of this group were eventually autopsied and cultures prepared from the spleen of each (8).

All tests were read at 24 hours; a reaction with 5 millimeters or more of induration was considered positive.

RESULTS

The results of these repeated skin tests on normal guinea pigs are summarized in tables 1-10.

Table 1.—Results of simultaneous tests on normal guinea pigs with 1-100 dilution of heat-killed yeast-phase antigen of Histoplasma capsulatum and with specified dilutions of histoplasmin, lots H-15 and H-6

	Heat- killed			Histor	lasmin				
	yeast- phase antigen	1	Lot H-15	5		Lot H-6			
. Item	Dilution								
	1-100	Undi- luted	1-10	1-100	Undi- luted	1-10	1-100		
Number of animals tested	10 0 0	10 0 0	10 0 0	10 0	10	10 0 0	10		

From table 1 it can be seen that none of the 10 animals which were first tested simultaneously with a skin test dose (0.1 ml.) of a 1–100 dilution of the heat-killed yeast-phase antigen of *Histoplasma* and the undiluted, 1–10, and 1–100 dilutions of histoplasmin, lots H–15 and H–6, reacted to any of these antigens in these dilutions. However, when retested 35 days later with the 1–100 dilution of the yeast-phase antigen and each of the three dilutions of lot H–15 histoplasmin (table 2) eight reacted to the undiluted H–15 but none to the other dilutions of H–15 nor to the yeast-phase antigen. Furthermore, on

Table 2.—Results of retesting guinea pigs with 1-100 dilution of heat-killed yeastphase antigen of Histoplasma capsulatum and with specified dilutions of histoplasmin, lot H-15.

	Heat- killed yeast-		Histor	olasmin			
	phase antigen		H-15				
Item		79 days					
	Dilution						
	1-100	Undiluted	1-10	1-100	1-100		
Number of animals tested	10 0 0	10 8 80. 0	10 0 0	10 0 0	9 2 22. 2		
Average diameter of reaction 1		8. 5			5, 5		

¹ Induration in millimeters.

the seventy-ninth day (table 2), two of nine, having received one test with undiluted H-6 and two with undiluted H-15 (tables 1 and 2), reacted to a 1-100 dilution of H-15. One of these two still reacted to this dilution of H-15 after 182 days.

Of the 10 animals which were tested simultaneously with a 1-10 dilution of the yeast-phase antigen and with undiluted, 1-10, and

Table 3.—Results of simultaneous tests on normal guinea pigs with 1-10 dilution of heat-killed yeast-phase antigen of Histoplasma capsulatum and with specified dilutions of histoplasmin, lots H-15 and H-6

	Heat- killed			Histor	lasmin				
Item	yeast- phase antigen		Lot H-15		Lot H-6				
tien	Dilution								
	1-10	Undi- luted	1-10	1-100	Undi- luted	1-10	1-100		
Number of animals tested	10 9	10	10	10	10	10	10		
Percentage of reactors	90, 0 6, 7	0	0	0	0	0	0		

¹ Induration in millimeters.

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22. 2 5. 5 1–100 dilutions of histoplasmin, lots H–15 and H–6 (table 3), 9 reacted on the first test to the 1–10 dilution of the yeast-phase antigen but not to the other antigens in the dilutions employed. When retested 35 days later with the same dilutions of H–15 and a 1–100 dilution of the yeast-phase antigen (table 4), 7 of 9 reacted to undiluted, 1 of 9 to the 1–10 dilution of H–15 and 4 of 9 to the 1–100 dilution of the yeast-phase antigen, but none to the 1–100 dilution of H–15. At 79 days, however, 6 of 9, having received 1 test with a 1–10 dilution of yeast-phase antigen, 2 with undiluted H–15 and 1

Table 4.—Results of retesting guinea pigs with 1-100 dilution of heat-killed yeast-phase antigen of Histoplasma capsulatum and with specified dilutions of histoplasmin, lot H-15

•	Heat- killed yeast- phase antigen			plasmin H-15			
Item	35 days after initial tests 79 c						
	1-100	Undiluted	1-10	1-100	1-100		
Number of animals tested	9 4 44. 4 5. 4	9 7 77.8 8.5	9 1 11. 1 5. 0	9 0 0	9 6 66. 7 6. 7		

Induration in millimeters.

with undiluted H-6, reacted to a 1-100 dilution of H-15. Two of these still reacted to this dilution of H-15 after 182 days.

Similar results were obtained with the 10 normal guinea pigs tested with undiluted blastomycin (tables 5 and 6), and with the undiluted

Table 5.—Results of simultaneous tests on normal guinea pigs with 1-100 dilution of heat-killed yeast-phase antigen of Blastomyces dermatitidis and with specified dilutions of blastomycin, lots B-7 and B-2

	Heat- killed yeast- phase		Lot B-7		omycin	Lot B-2		
ltem	Dilution					Lot B-2		
	1-100	Undi- luted	1-10	1-100	Undi- luted	1-10	1-100	
Number of animals tested Number of reactors Percentage of reactors A verage diameter of reaction ¹	11 1 9.1 5.0	11 2 18. 2 5. 0	11 0 0	11 0 0	11 0 0	11 0 0	11 0 0	

Induration in millimeters.

Table 6.—Results of relesting guinea pigs with 1-100 dilution of heat-killed yeast-phase antigen of Blastomyces dermatitidis and with specified dilutions of blast-omycin, lot B-7

	Heat- killed yeast- phase anti- gen Lot B-7						
Item		79 days					
	1-100	Undiluted	1-10	1-100	1-100		
Number of animals tested	10	10	10	10	10		
Percentage of reactors. A verage diameter of reaction i	0	90. 0 7. 3	0	0	10. (5. (

Induration in millimeters.

Table 7.—Results of testing normal guinea pigs with 1-10 dilution of autoclaved filtrate of Candida (Monilia) albicans and of retesting 15 days later with 1-10 and 1-100 dilutions of the filtrate

*	Autoclaved filtrate of C. albicans					
	Day of observation					
ltem	1	15				
	Dilution					
	1-10	1-10	1-100			
Number of animals tested Number of reactors Percentage of reactors Average diameter of reactions 1	11 0 0	11 5 45. 5 5. 3	11 3 27.3 5.7			

¹ Induration in millimeters.

and 1-10 dilutions of the autoclaved filtrate antigen prepared from *C. albicans* (tables 7 and 8). Repeated testing with a 1-100 dilution of the autoclaved filtrate of *C. albicans*, however, did not result in any reactions to this dilution of this antigen on the second test 15 days after the first.

Table 8.—Results of testing normal guinea pigs with undiluted autoclaved filtrate of Candida (Monilia) albicans and of retesting 15 days later with undiluted and with 1-10 and 1-100 dilutions of the filtrate

Item	Autoclaved filtrate of C. albicans Day of observation						
		Dilution					
	Undiluted	Undiluted	1-10	1-100			
Number of animals tested	12 0 0	12 6 50.0 7.8	12 2 16.7 5.0	12 1 8.3 5.0			

Induration in millimeters.

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Table 9.—Results of simultaneous tests on normal guinea pigs with 1-100 dilution of heat-killed antigen of Candida (Monilia) albicans and with 1-100 dilution of histoplasmin, lot H-15

to a programme of the second	Heat-killed antigen C. albicans	Histoplasmin Lot H-15	
Item	Dilution		
•	1-100	1-100	
Number of animals tested Number of reactors Percentage of reactors Average diameter of reaction ¹	20 9 45.0 5.7	20 0 0	

¹ Induration in millimeters.

Table 10.—Results of retesting guinea pigs with heat-killed antigen and with autoclaved filtrate of Candida (Monilia) albicans and with histoplasmin, lot H-15

	Heat-killed antigen C. albicans	Autoclaved filtrate of C. albicans	Histoplasmin Lot H-15		
The state of the s	Day of observation				
Item	15	1	82		
		Dilution	The second		
	1-100	1-100	1-100		
Number of animals tested Number of reactors. Percentage of reactors. Average diameter of reaction 1	20 20 100.0 7.9	19 16 84. 2 7. 1	19 12 63. 2 8. 5		

Induration in millimeters.

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Of the 20 normal animals tested with the heat-killed suspension of *C. albicans*, 9 reacted to the first injection of a 1-100 dilution (table 9) and 20 of 20 to the same amount of the same antigen when retested 15 days later (table 10).

Cultures were prepared from the spleen of each of 11 guinea pigs in the first group, tested repeatedly with histoplasmin; 4 in the second group, tested with blastomycin; and 15 from the fourth group tested with the heat-killed suspension of *C. albicans*. All were negative for pathogenic fungi as reported elsewhere (8).

DISCUSSION

It has been shown previously (1, 2) that repeated testing with a 1–100 dilution of various lots of histoplasmin and blastomycin and heat-killed antigens prepared from the yeast-phase of *Histoplasma capsulatum* and *Blastomyces dermatitidis* does not irritate nor sensitize normal guinea pigs. However, from the evidence presented above (tables 1–6) it would seem that repeated skin tests with 0.1 ml. doses of undiluted histoplasmin or blastomycin, or two or more injections (intradermally) of the undiluted antigens simultaneously, or tests with a 1–10 dilution of the heat-killed yeast-phase antigen prepared from cultures of *Histoplasma* may sensitize normal guinea pigs to some degree so that they may subsequently react to tests with a 1–10 or occasionally a 1–100 dilution of the same antigen whereas they did not react to these dilutions previously.

The finding that multiple tests with undiluted histoplasmin or blastomycin may sensitize is, as indicated above, based in part on work previously reported (1, 2). In these papers it has been shown, for example, that repeated tests with 1–10 and 1–100 dilutions of these particular lots of histoplasmin and blastomycin do not sensitize. Therefore, it is assumed that when used simultaneously with the undiluted antigens, any subsequent reactions of these animals to these antigens are due to the previous use of the undiluted materials.

Similar results have been obtained with an autoclaved filtrate, which is probably carbohydrate in nature, prepared from Candida

(Monilia) atbicans (tables 7 and 8).

The finding that these filtrate antigens, if used in sufficient concentration, may sensitize normal guinea pigs to some degree is supported by the observations of Hirsch and D'Andrea (3) and T'Ung and Wong (5). The former found that normal guinea pigs could be sensitized by repeated subcutaneous injections of coccidioidin so that they would subsequently react to an intradermal injection of coccidioidin or to the "specific substance," probably carbohydrate in nature, which was isolated from coccidioidin. T'Ung and Wong found that a polysaccharide isolated from Monilia (Candida) tropicalis, injected intraperitoneally, was capable of inducing a low degree of sensitivity in guinea pigs, although this substance was inferior to the whole yeast cells as a sensitizing agent.

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In table 9 it can be seen that 9 of 20 normal guinea pigs reacted to a skin-test dose of a 1-100 dilution of a heat-killed suspension of Candida (Monilia) albicans on the first injection. Fifteen days later, 20 of 20 reacted to a second injection of the same amount (table 10). Several investigators have found that in animals, injection of whole, dead cells of Candida usually sensitizes (4, 6) although such sentitization is weaker than that produced by actual invasion of the tissues by living cells.

The subsequent reaction of the group of animals sensitized by intradermal injections of the heat-killed suspension of C. albicans (table 10) to a 1-100 dilution of lot H-15 histoplasmin must be regarded as evidence of a cross reaction, although it should be pointed out that this dilution of this lot of histoplasmin is approximately 10 times the critical titer of this antigen for guinea pigs (2). other animals sensitized with living cells of C. albicans have shown a much lower degree of cross reaction with this lot of histoplasmin (9).

The sensitization of normal animals by repeated injections or two or more simultaneous injections of undiluted histoplasmin or by the use of a 1-100 heat-killed suspension of C. albicans would seem to explain the reactions of normal animals to histoplasmin reported in the following paper (8).

SUMMARY

The effects of simultaneous and repeated intradermal injections of 0.1 ml. of various dilutions of histoplasmin, blastomycin, an autoclaved filtrate of Candida (Monilia) albicans, heat-killed suspensions of the yeast-phase of Histoplasma capsulatum and Blastomyces dermatitidis, and of a 1-100 dilution of a heat-killed suspension of C. albicans on normal guinea pigs have been studied.

It has been shown that these antigens, if used in sufficient concentrations, may sensitize normal guinea pigs.

REFERENCES

- (1) Emmons, C. W., Olson, B. J., and Eldridge, W. W.: Studies of the role of fungi in pulmonary disease. I. Cross reactions of histoplasmin.
- Health Rep. 60: 1383 (1945).

 (2) Howell, Arden: Studies of fungus antigens. I. Quantitative studies of crossreactions between histoplasmin and blastomycin in guinea pigs. Health Rep. 62: 631 (1947).
- (3) Hirsch, E. F., and D'Andrea, D.: Sensitization of guinea pigs with broth culture filtrates and with killed mycelium of Coccidioides immitis. J. Inf. Dis. 40: 638 (1927).
- (4) Kurotchkin, T. J., and Lim, C. E.: Anaphylaxis with water soluble specific substance from yeast-like fungi. Proc. Soc. Exper. Biol. and Med. 28:
- (5) T'Ung, T., and Wong, S. C.: Sensitizing capacity of polysaccharide of Monilia tropicalis. Proc. Soc. Exper. Biol. and Med. 41: 155 (1939).
 (6) Lim, C. E., and Kurotchkin, T. J.: Quoted in: Skinner, C. E.: The yeast-like fungi: Candida and Brettanomyces. Bact. Rev. 11: 227 (1947).
 (7) Martin, D. S. Schallenger the immunicación relationships among various.
- (7) Martin, D. S.: Studies on the immunologic relationships among various
- species of the genus Candida (Monilia). Am. J. Trop. Med. 22:295 (1942).

 (8) Howell, Arden: Isolation of pathogenic fungi from experimentally inoculated guinea pigs. This issue, page 602.

 (9) Howell, Arden: Unpublished data.

ISOLATION OF PATHOGENIC FUNGI FROM EXPERIMEN-TALLY INOCULATED GUINEA PIGS ¹

By Arden Howell, Jr., Ph. D., Senior Mycologist, Public Health Service

INTRODUCTION

Numerous investigators have reported variable results in attempts to infect laboratory animals with *Histoplasma capsulatum* (1–8), *Blastomyces dermatitidis* (9–16), and *Candida albicans* (17–21). The following study was undertaken, therefore, to determine (1) a sublethal dosage of each of these fungi which would sensitize the animals to skin test antigens prepared from the homologous fungi and (2) to learn if these fungi could be isolated from the inoculated animals after variable periods of time, with or without the production of an apparent generalized infection.

MATERIALS AND METHODS

The strains of *H. capsulatum* (22) and *B. dermatitidis* (23) employed in these studies were those previously reported. The strains of *Candida* (*Monilia*) albicans were isolated from specimens received at the United States Public Health Service laboratory at the University of Kansas Hospital.

Each of 80 normal albino guinea pigs was inoculated intraperitoneally with graded doses of a 1–100 saline suspension of the living yeast phase of *H. capsulatum*, 82 with graded doses of a similar suspension of the living yeast phase of *B. dermatitidis*, and 35 with graded doses of a 1–100 saline suspension of living cells of *Candida* (*Monilia*) albicans.

The suspension of the yeast phase of *Histoplasma* was prepared from 5-day-old cultures grown on sealed blood agar slants and incubated at 37° C. (24); that of *Blastomyces* from 7-day-old cultures of the yeast phase grown on Difco brain heart infusion agar at 37° C.; and that of *C. albicans* from 2-day-old cultures grown on Sabouraud's agar at 37° C.

The dosages employed are shown in table 1.

Table 1.—Number of guinea pigs inoculated intraperitoneally with specified dosages of 1-100 suspension of yeast-phase of Histoplasma capsulatum, of Blastomyces dermatitidis, and of Candida (Monilia) albicans

		Dosage (ml. 1-100 suspension per gram of body weight)									
	0. 00100- 0. 00199	0, 00200- 0, 00299					0, 00700- 0, 00799		0, 00900- 0, 00999	Total	
Histoplasma capsula- tum Blastomyces dermati-	7	13	20	16	15	9	0	0	0	86	
tidisCandida albicans	5 7	9 12	19 12	26 4	8	4 0	5 0	5 0	1 0	82 33	

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All animals were skin-tested prior to inoculation with a skin-test dose of a 1-100 dilution of an antigen prepared from the homologous organism [i. e., histoplasmin, blastomycin, or an antigen prepared from *C. albicans* (25)]. No animal reacted to any antigen in the dilution employed.

The animals were weighed at the time of inoculation and at intervals thereafter and were skin-tested with various antigens at intervals. Thirteen of those inoculated with *Histoplasma* died between 30 and 247 days after inoculation; the remainder were sacrificed at intervals varying from 2 to 10 months. Twenty-eight of those inoculated with *Blastomyces* died between 18 and 289 days after inoculation; the remainder were sacrificed at intervals of 1 to 11 months. Four of the group inoculated with *C. albicans* died between 11 and 63 days after inoculation; 24 of the remainder were sacrificed at intervals varying from 49 to 76 days after inoculation.

All animals were autopsied and any marked pathological change was noted. If abscesses or other lesions were present material from these lesions was used for culture. In all animals in which there were no marked pathological changes the spleen was removed, a portion of each fixed for sectioning, and the remainder used for cultures.

Cultures from the animals inoculated with *Histoplasma* were made on each of two plates of brain heart infusion blood agar and potato dextrose agar as previously reported (22); those from animals inoculated with *Blastomyces* or *C. albicans* on two plates of the same blood agar and two plates of Sabouraud's agar. Streptomycin and penicillin in concentrations of 40 and 20 units, respectively, per milliliter of medium were added to all media. One plate of each medium was then incubated at 37° C. and the other at room temperature (approximately 25° C.).

In addition to the animals which were inoculated with *Histoplasma*, *Blastomyces*, and *C. albicans*, 41 normal albino guinea pigs were studied as controls. These animals were divided into three groups. The first group, composed of 10 animals, were skin-tested with a skin-test dose of a 1–100 dilution of each of several antigens on the day they were received at the laboratory and sacrificed 3 days later. The second group, composed of 15 animals, were kept under observation for a total of 182 days and were kept in the normal animal room. The third group, composed of 16 animals, were kept in the same room with the inoculated animals described above and were under observation for 232 to 259 days. Three of this group died between the 244th and 258th day of observation. All of the animals in groups two and three were skin tested at intervals with various antigens. All were then sacrificed at the end of the periods indicated.

Each of these 41 normal animals was autopsied. At autopsy any

marked pathological changes were noted and the spleen of each removed. A portion of the spleen of each was fixed for sectioning and the remainder used for cultures.

Cultures of each were made on two plates of the brain heart infusion blood agar, one plate of potato dextrose agar and one plate of Sabouraud's agar. One plate of the blood agar was incubated at 37° C.; the other three plates at room temperature (approximately 25° C.).

All positive cultures or cultures suspected of being positive were confirmed by subculturing the suspected colonies on the appropriate medium at room temperature.

RESULTS

1. Animals inoculated with Histoplasma capsulatum.—With the dosages employed, in animals inoculated intraperitoneally with the living yeast phase of H. capsulatum there appeared to be no correlation between the size of the inoculating dose and the number of animals which developed an apparently generalized infection.

Of the 80 guinea pigs in this group, an apparently generalized infection developed in 3, or 3.8 percent, of the total inoculated (table 2).

Table 2.—Summary of findings on guinea pigs developing apparently generalized infection following intraperitoneal inoculation with 1-100 saline suspension of yeast-phase of Histoplasma capsulatum

			Skin re	action to			weight ange			-3
Guinea	Dosage (ml. 1-100 suspen- sion per		plasmin, H-15	fractio	echaride on (his- asmin, H-17)			Time	Gross pathology	Results of culture
pig No.	gram of body weight)	1-100	dilution	1.0 1	ng./ml.	Gms.	Per- cent- age	death (Days)		results of culture
		I	Days after	inoculat	ion					
		40	75	89	47					
230 260 267	0. 00345 0. 00294 0. 00291	=	_	-	+,	-110 +15 -185	-25. 3 +2. 9 -35. 9	1 90 48 3 43	Moderate . Moderate . Moderate .	Lung+; spleen+ Spleen+ Spleen unknown (contaminated)

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1 Sacrificed.

Only 5 mm. of induration.
 Died as a result of a laboratory accident.

Two of these three animals underwent a progressive loss of weight, the third showed a slight increase in weight at the time of death. Of the two which showed a progressive weight loss, one died on the forty-third day apparently from a laboratory accident and the other was sacrificed 90 days after inoculation. None of the three reacted to any skin test at any time except one (table 2). All three showed moderate pathology at autopsy, particularly in the thoracic cavity.

Cultures prepared from the lungs of guinea pig No. 230 were positive for *Histoplasma*: cultures of the lungs of the other two animals were overgrown with bacteria. Cultures prepared from the spleens of guinea pigs Nos. 230 and 260 were positive; cultures of the spleen of the third were overgrown with contaminants.

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One of the 80 apparently was neither infected not sensitized by the inoculation. Repeated skin tests with a 1-100 dilution of histoplasmin, lot H-15 (23) were all negative. No gross pathology was noted at autopsy 91 days after inoculation, by which time the animal had gained approximately 20.0 percent in weight. Sections and cultures made from the spleen at autopsy were both negative for Histoplasma.

Of the remaining 76 animals inoculated with *Histoplasma*, 11 died (table 3). Two of these apparently died from laboratory accidents. These two reacted to a skin-test dose of a 1-1000 dilution of histoplasmin, lot H-15, 26 days after inoculation; both showed a marked increase in weight; neither showed any marked pathology at autopsy. However, cultures prepared from the spleens of both were positive.

The cause of death of the other 9 animals which died during the course of the experiment was not determined. All reacted to a skintest dose of a 1-1000 dilution of histoplasmin, lot H-15, 26 days after inoculation, and 2 to a 1-100 dilution of this same antigen 202 days after inoculation. Eight of the nine gained a considerable amount of weight while under observation; only two showed any marked pathology at autopsy. In these two, guinea pigs Nos. 193 and 200, there were slight to moderate changes in the thoracic cavity. Cultures of material from the lungs of both and of the spleen of No. 193 were overgrown with bacteria. Cultures of the spleen of guinea pig No. 200 yielded Histoplasma. Of the remaining seven which showed essentially no pathology at autopsy, Histoplasma was isolated from the spleens of four. These 4 animals were autopsied 79, 122, 30, and 140 days after inoculation. There is no clear evidence, therefore, that any of these 11 animals died from experimental histoplasmosis.

The remaining 65 guinea pigs remained apparently healthy. These were sacrificed 2 to 9 months after inoculation. All, at some time, reacted to a skin-test dose of at least a 1-100 dilution of histoplasmin, lot H-15, and 54 reacted, at some time, to a 1-1000 dilution of this antigen. All gained a considerable amount of weight, and none showed any marked pathology at autopsy. However, *Histoplasma* was isolated from the spleens of 39, or 60.0 percent of these animals. Among 31 animals kept under observation for a period of 6 months or longer, cultures prepared from 14, or 45.2 percent, were positive. It is interesting that those animals from which cultures were obtained were not necessarily those which had received the greater doses.

not necessarily those which had received the greater doses.

It was possible to isolate Histoplasma from the spleens of 4 of 10

Table 3.—Summary of findings on guinea pigs dying 1 after inoculation with 1-100 suspension of yeast-phase of Histoplasma capsulatum

Guinea pig No. gram of body weight)	9	Dilution			Mynt seedles	of alaman			
	- 0	4744444	n(Net weight change	nt change			
Weight) 0.00164 0.00333 0.00267 0.00283			1-100				Time of death (days)	Gross pathology at autopsy	Results of culture
0, 00164 0, 00333 0, 00267 0, 00263		Days after inoculation	oculation		Gms.	Percent- age			
0,00164 0,0033 0,00267 0,00263	_	171 186	202	228					
0.00267					+470	+154.0	163	Essentially none	Lung -; spleen Spleen +.
000000		8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8	E E E E E E E E E E E E E E E E E E E	1 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	++580	+74.7	3 3	Essentially none	Spleen +.
1		+	+	6	+1455	+102.3	300	Essentially none Essentially none Essentially none	Spleen +.
* * * * * * * * * * * * * * * * * * *	1 1			8	+180	+40.0	106	Essentially none	Spleen +. Spleen unknown (contaminated);
0.00441 + 200 0.00455 +		+	+	+	+310	+91.2	247	Essentially none	Spleen + lung unknown (con-

 1 Exclusive of guinea pigs number 260 and 267 included in table 1. 2 Died as a result of laboratory accidents.

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apparently normal, healthy animals which had been inoculated with large doses of this fungus intraperitoneally at least 9 months previously.

None of these 65 animals, at any time during the course of the experiment, nor at autopsy, showed any signs of infection with the fungus except for the development of a sensitivity to the homologous organism

or its products at some time after inoculation.

Of the 77 animals which failed to develop any visible signs of experimental histoplasmosis, other than a skin sensitivity to histoplasmin, it was possible to isolate *Histoplasma*, after variable periods of time, from the spleens of 46, or 59.6 percent.

2. Animals inoculated with Blastomyces dermatitidis.—In animals inoculated intraperitoneally with the living yeast phase of Blastomyces dermatitidis, as with the animals similarly inoculated with Histoplasma, there appeared to be no correlation, with the narrow ranges of dosages employed, between the size of the inoculating dose and the number of animals which developed an apparent generalized infection

Of 82 guinea pigs in this group, 17 developed an apparent infection (table 4); 13 of these 17 died between the 18th and 127th day after inoculation; the remaining 4 were sacrificed between the 41st and 257th day. Of the 13 which died, 11 showed moderate to marked loss of weight, 5 consistently failed to react to a skin-test dose of a 1–100 dilution of blastomycin, lot B–2 (23); 10 showed extensive, and 3 moderately extensive, pathology at autopsy; cultures obtained from one or more organs of each were positive for Blastomyces. One of these, guinea pig No. 117, died as a direct result of a laboratory accident. This animal had failed to react to a skin test dose of a 1–100 dilution of blastomycin, lot B–2, and even though it had gained a considerable amount of weight, showed extensive pathology at autopsy.

Of the 4 in the above group which were sacrificed, 3 showed a slight to marked increase in weight; all 4 failed repeatedly to react to a skintest dose of a 1-100 dilution of blastomycin, lot B-2 or B-7 (23); and 2 showed moderate to extensive pathology at autopsy. A third, sacrificed 257 days after inoculation, by which time it had gained over 200 grams, or 47.9 percent, in weight, showed an abscess in the liver approximately 5 cm. in diameter at autopsy. Cultures obtained from one or more tissues of each of these 4 animals were positive for Blastomyces. The essential findings on each of these 17 guinea pigs are summarized in table 4.

Of the remaining 65 guinea pigs, 15 died (table 5). Two of these apparently died as a result of laboratory accidents. One of these reacted to a skin-test dose of a 1-1000 dilution of blastomycin, lot B-2, 47 days after inoculation, while the other did not. Both gained

Table 4.—Summary of findings on guinea pigs developing apparently generalized infection following intraperitoneal inoculation with 1-100 saline suspension of yeast-phase of Blastomyces dermatitidis

			skin reactiv	Skin reaction to blastomycin	myein		Net weight change	it change			
		Lot	Lot B-2	I	Lot B-7						
	Dosage (ml. 1-100 suspen-			Dilution		-					
Guines pig No.	sion per gram of body weight)	1-1000	1-100	1-1000	1-100	90	Gms.	Percen- tage	death (days)	Gross pathology at autopsy	Results of culture
			Days aft	Days after inoculation	00						
			47								
113 116 117 122 123 131 131	0.00471 0.00158 0.00440 0.00450 0.00240 0.00198 0.00198	+	1 1111	6			-210 -25 -128 +139 -150 -100	-33.0 -7.9 -7.9 +128.5 -1.9	28 8 18 8 18 8 18 8 18 8 18 8 18 8 18 8	Extensive Extensive Extensive Extensive Extensive Extensive Extensive	Liver +; spleen +; lung +. "Pancreas" +. "Pancreas" +; spleen +. "Pancreas" +; spleen +. Liver abscess +. Spleen +; subcut, absc. +. Spleen +.
			28		187	252					
147	0.00566 0.00670 0.00799		111		1	1	-220 -160 +210	-31.1 -26.8 +47.9	1 190 40	Mcderate Extensive Liver abscess.	Spleen +; "pancreas" +. Spleen +; liver absc. +. Spleen +; liver absc. +.
			4	92		40					
233 239 262 270 271	0. 00364 0. 00284 0. 00391 0. 00531		+		* 1777		135	- 24.5 - 25.4 - 25.9 - 45.9 - 45.9	113 93 52 70	Extensive. Liver abscesses. Liver abscesses. Moderate. Extensive	Spleen +; abdom. absc. +. Liver abscess +. Liver abscess +. Liver abscess +. Liver abscess +.
			38	64							
132	0.00667		188	+		1 6 8 1 0 1 0 1 0 1 0 1 0 1 0 1 0 1 0 1 0 1	-45	-7.5	127	Extensive	Subcut, abscess +.
137	0.00449		1			0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	+40	+9.0		141 Moderate	Spleen +; "pancreas" +.

Weight at 56 days.

[.] Died as a result of laboratory accident.

a large amount of weight before death. At autopsy, both showed some degree of pathology. Cultures from material taken at autopsy of both animals were negative for *Blastomyces*, though those from one of the animals were overgrown with bacteria.

The cause of death of the other 13 animals which died was undetermined. Twelve showed a marked increase in weight; and 10 at some time reacted to one or more repeated skin tests with a 1–1000 dilution of blastomycin, lot B–2 or B–7. At autopsy, six showed some degree of pathology. Of these, only one gave cultures positive for *Blastomyces*, while of the seven which showed essentially no pathology, *Blastomyces* was isolated from the spleen of two and the lungs of one. It would seem doubtful, therefore, that any of these animals died from experimental blastomycosis. The essential findings on each of these 15 animals are shown in table 5.

The remaining 50 guinea pigs remained apparently healthy. All, at some time, reacted to a skin-test dose of at least a 1-100 dilution of blastomycin, lot B-2 or B-7; 22 at some time to at least a 1-1000 dilution of lot B-2, and 37 to at least a 1-1000 dilution of lot B-7. All gained markedly in weight. None showed any marked pathology at autopsy, although there were small abscesses in the liver of two, in the pelvic region of two, "pancreas" of one, and abdominal wall of one. However, Blastomyces was isolated from 1 or more tissues removed at autopsy from 26, or 52.0 percent, of these animals. Among 33 animals kept under observation for a period of 6 months or longer cultures prepared from 15, or 45.5 percent, were positive.

It was possible to isolate *Blastomyces dermatitidis* from the spleens of four of 9 guinea pigs which had been inoculated intraperitoneally with this fungus at least 10 months previously.

Of the 65 which failed to show any visible signs of experimental blastomycosis, other than a skin sensitivity to blastomycin, it was possible to isolate *Blastomyces*, after variable periods of time, from 30, or 46.2 percent.

As with the animals inoculated with *Histoplasma*, the animals from which cultures were obtained were not necessarily those which received the greater doses.

3. Animals inoculated with Candida (Monilia) albicans.—Of 35 guinea pigs inoculated with C. albicans, none developed an apparently generalized infection. Four died between the 11th and 63d day after inoculation (table 6). The animal which died on the 11th day apparently died from an intestinal obstruction. The other three reacted to at least a 1-100 dilution of an autoclaved filtrate antigen prepared from the homologous fungus (25), two of them to a 1-1000 dilution, on the 41st day. Two of the three lost weight. However, there were no marked gross pathological findings at autopsy in any one of the four, and cultures from two were negative for C. albicans. No

Table 5.—Summary of findings on guinea pigs dying 1 after inoculation with 1-100 suspension of yeast-phase of Blastomyces dermatitidis

			30	kin reacti	Skin reaction to blastomycin	omycin			Net weight change	t change			
			Lot B-2	-2		Lot B-7	1-					1/2	
on aire	Dosage (ml. 1-100 suspen-				Dilution						Time of	Gross pathology at	Besulte of online
umea pig no.	gram of body weight)	1-1000	000	1-100	1-1000		1-100		Gms.	Percent-	(days)	autopsy	arian or carrier
				Days at	Days after inoculation	ition							
		47	25		25	215	250	280					
101 002 003 221 226 236 237 24 24 256 256 27 27 27 27 27 27 28	0,00447 0,00428 0,00330 0,00151 0,00151 0,00412 0,00412 0,00412 0,00425 0,00459	+++1++++1	++1 +++ +		+++ +++ +	+ ++	++		+110 +270 +195 +1165 +110 +110 +110	++++++ 22.50 1.7.28 2.20 2.7.30 2.7.30	227 208 170 170 289 289 289 288 1153 163 8	Essentially none Essentially none Moderate Moderate Necrosis of liver Liver absects Siller (tungs) Siller (tungs) Siller (tungs)	Lung
				Days	Days after inoculation	lation							
				88	88								
149 157 162	0.00860 0.00785 0.00802			+11	+				+150	+ 53.8	158 138	Absc. in diaphragm Essentially none	Spleen – . Spleen – . Spleen – .
* * * * * * * * * * * * * * * * * * *	0.00769			Days a	Days after inoculation	tion					186	Essentially none	Lung +.
					35	75	98	125					
201	0.00400				+	1	+	+	+155	+31.0	130	Moderate	Spleen +.

In addition to those summarized in table 4.

Died as a result of laboratory accidents.

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Table 6.—Summary of findings on guinea pigs dying after inoculation with 1-100 suspension of Candida (Monilia) albicans

Guinea pig num-	Dosage (ml. 1-100 suspen- sion per gram	to auto	after		veight ange	Time of death (days)	Gross pathology	Results of culture
ber	of body wgt.)	Dilu	tion	Gms.	Per-	(days)		
	- 11	1-1000	1-100	Cins.	age			
351	0. 00351	+	+	+40	+7.0	63	Post mortem	No culture made
357	0.00431	-	+	-90	1-15.5	54	Essentially none	Spleen
368	0. 00364	+	‡	-100	-18.2	45	Abdominal adhesions.	Spleen-; lung-;
383	0.00235			-30	-7.1	11	Intestinal obstruc- tion.	No culture made

Weight taken 5 days before death.

cultures were made from the other two. Therefore, although the cause of death of three of these animals was not determined, it would not seem that they died as a result of the inoculation of *C. albicans*.

Of the remaining 31 animals, 24 were sacrificed. All of these showed a definite and constant increase in weight, although the group sacrificed during the third month were markedly dehydrated in appearance at the time they were sacrificed. All but one reacted to a skin-test dose of at least a 1–100 dilution of the autoclaved filtrate antigen of C. albicans 41 days after inoculation. At autopsy, none of the 24 which were sacrificed showed any gross pathology except for abdominal adhesions in five. Cultures prepared from the spleens of each of these 24 were negative except for one sacrificed on the 49th day, from which C. albicans was isolated. It would seem, therefore, that none of these animals developed a widespread experimental infection, with the dosages employed for inoculation, although the majority (33 of 35, or 94.3 percent) did develop a skin sensitivity to the homologous fungus or its products.

4. Normal animals.—Forty-one normal guinea pigs, divided into 3 groups, were included in this study as controls.

The first group, composed of 10 animals, were tested with a skintest dose of a 1-100 dilution of histoplasmin, lot H-15, blastomycin, lot B-7, and the autoclaved filtrate antigen of *C. albicans* on the same day the animals were received into the laboratory. None reacted to any of these antigens in this dilution. Each of these animals was sacrificed 3 days later. At autopsy none showed any gross pathology; cultures from the spleen of each were negative for pathogenic fungi.

The second group, composed of 15 animals, were kept under observation for a period of 182 days in the normal animal room. At the time they were sacrificed (on the 182d day of observation) 4 reacted to a skin-test dose of a 1-100 dilution of histoplasmin, lot

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H-15.² None reacted to the same dilution of blastomycin, lot B-7. At autopsy none showed any gross pathology; and cultures from the spleen of each were negative for pathogenic fungi.

The third group, composed of 16 animals, were kept in the same room with the animals inoculated with *Histoplasma*, *Blastomyces*, and *C. albicans*. Three of these animals died between the 244th and 258th day of observation. At autopsy two showed some gross pathology, but cultures from the spleens of each of the three were negative for

pathogenic fungi.

Of the remaining 13, 8 were sacrificed on the 232d day of observation. At this time, three of these reacted to a skin-test dose of a 1–100 dilution of histoplasmin, lot H–15, and four to the same dilution of the *C. albicans* antigen.² None, however, reacted to a 1–100 dilution of blastomycin, lot B–7. The remaining 5 were sacrificed on the 259th day. At this time none of these reacted to a skin-test dose of a 1–100 dilution of histoplasmin, lot H–15, blastomycin, lot B–7, or the autoclaved filtrate antigen prepared from *C. albicans*. None of these 13 animals showed any marked gross pathology at autopsy. Cultures from the spleens of each were negative for pathogenic fungi.

DISCUSSION

From the data presented above it would seem that guinea pigs are relatively resistant to infection with H. capsulatum when this fungus is inoculated intraperitoneally, and that, in the dosage range used, there was little correlation between the dosage employed for inoculation and the number which developed a generalized extensive infection. Only 3 of 80, or 3.8 percent, of the animals included in this study appeared to develop such an infection from an intraperitoneal inocu lation of relatively large doses of the living yeast phase from 5-day-old cultures incubated at 37°C. This finding is in general agreement with most of the reports on infection of these animals with this fungus. Reid et al. (5), for example, reported a generalized infection in one guinea pig inoculated with growth from a culture of the yeast Hansmann and Schencken (2) reported that guinea pigs inoculated intraperitoneally with material from a biopsy of skin and lymph nodes from a human case of histoplasmosis, from which Histoplasma was subsequently isolated in culture, failed to show any evidence of infection several months after inoculation. Similarly, Parsons and Zarafonetis (8) failed to induce infection in guinea pigs with material from a biopsy from a human case.

Of the remaining 77 animals in the series reported above (table 1), 76 developed some degree of skin sensitivity. This is evident from the fact that all of these 76 reacted, at some time after inoculation,

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² These reactions are explained in Studies of Fungus Antigens, III. See p. 595.

to a skin-test dose of at least a 1-100 dilution of histoplasmin, lot H-15, whereas none of them had reacted to this dilution of this antigen prior to inoculation.

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It has also been demonstrated that *Histoplasma* may be recovered from the spleens of apparently healthy guinea pigs inoculated with this fungus as long as 10 months previously although the animals showed no signs of gross pathology at autopsy. This is in agreement with a recent paper by Emmons et al. (26) in which was reported the isolation of *Histoplasma* from 1 mouse and 10 rats, none of which showed any gross lesions at autopsy. From these animals *Histoplasma* was isolated most frequently from the spleen and liver. Emmons (26) further reported that guinea pigs commonly recover from the acute phase of infection with *Histoplasma* and survive indefinitely and that it was possible to isolate *Histoplasma* regularly from experimentally infected guinea pigs kept in the laboratory under observation for over 2 years, even though there were no gross abnormalities observed at autopsy.

It would also seem that guinea pigs are relatively resistant to infection with B. dermatitidis, when this fungus is inoculated intraperitoneally, and that there is little correlation between the dosage employed and the number which develop a generalized infection. Only 17 of 82, or 20.7 percent, apparently developed such an infection from the intraperitoneal inoculation of large doses of the living yeast phase from a 7-day-old culture incubated at 37° C. Similar results have been reported by various investigators. Benham (15) reported that dogs and monkeys are very susceptible to infection with this fungus but that less extensive lesions develop in guinea pigs. DeMonbreun (16) found that guinea pigs inoculated with this fungus develop localized abscesses which soon healed following subcutaneous or intradermal inoculations. Others (9-14) have obtained variable results.

Although it would seem that guinea pigs are relatively resistant to an intraperitoneal inoculation of B. dermatitidis, a comparison of this group and the group inoculated with Histoplasma indicates that they are more susceptible to inoculation with this fungus than to inoculation with Histoplasma. For example, of 70 guinea pigs which received doses of Blastomyces comparable in volume of packed cells, to the 80 inoculated with Histoplasma, 16 of the 70 developed a generalized infection whereas only 3 of 80 of those inoculated with Histoplasma appeared to develop such an infection.

Of the remaining 65 guinea pigs inoculated with *Blastomyces*, 61 reacted, at some time after inoculation, to a skin-test dose of at least a 1-100 dilution of blastomycin, lot B-2 or B-7. None had reacted to this dilution of blastomycin prior to inoculation. The reason for the failure of the remaining four animals to become sensitized was not determined.

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It has also been demonstrated above that *Blastomyces* may be recovered from the spleens of apparently healthy guinea pigs which had been inoculated with this fungus as long as eleven months previously,

and which showed no gross lesions at autopsy.

Of 35 guinea pigs similarly inoculated with large doses of Candida (Monilia) albicans none apparently developed more than a transient infection. This is in accord with the results obtained by Ninni and Fittipaldi (21) who reported that one-fifth of an agar slant culture of pathogenic monilias inoculated intraperitoneally failed to kill guinea pigs, and of Benham (20) and Nye et al (19). Benham reported that guinea pigs are less susceptible to infection with C. (M.) albicans than are rabbits. Nye et al. reported that of 38 guinea pigs inoculated intraperitoneally with 250×10^6 organisms from 4-day broth cultures of Parasaccharomyces A. (pathogenic for rabbits) only one developed an extensive infection. This animal died on the fourth day after inoculation with a purulent peritonitis. Others (17, 18) have reported variable results.

Although none of the animals inoculated with C. albicans developed an apparently generalized infection, 33 of 35 were sensitized so that at 41 days after inoculation they reacted to a skin-test dose of at least a 1-100 dilution of an autoclaved filtrate antigen prepared from this fungus (25). This group of animals were similar to those inoculated with both Histoplasma and Blastomyces in that in the majority of the animals there were no demonstrable lesions at autopsy. However, they differed in that C. albicans was recovered in culture from the spleen of only one of 26 animals autopsied, even though all of the 26 died or were sacrificed not later than 76 days after inocu-

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lation.

Finally, it has been shown that none of these pathogenic fungi could be isolated from the spleens of 41 normal animals, 31 of which were kept under observation for a minimum of 182 days in the laboratory before autopsy. Sixteen of these normal guinea pigs were kept in the same room with the animals inoculated with *Histoplasma*, *Blastomyces*, and *C. albicans*. This would seem to indicate that these infections are not readily transmitted from one individual to another, and is in agreement with the report of Emmons et al. (26) that histoplasmosis was not transmitted from a naturally acquired infection in a dog to other dogs kept for months in the same cage with it. This failure to isolate a pathogenic fungus from these normal guinea pigs lends support to the conclusion that the cultures isolated from the incculated animals were isolated as a result of the inoculation.

It should be noted that although there appeared to be little correlation between the size of the dosage used and the number of animals which developed extensive infections with any of the three fungi studied, the dosages employed covered only a five-to nine-fold range.

It is possible that had larger doses been used for inoculation, some correlation might have become apparent. It would also seem that the percentage of experimentally inoculated animals from which H. capsulatum and B. dermatitidis can be isolated decreases with the length of time after inoculation.

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SUMMARY

A study has been made on the susceptibility of guinea pigs to intraperitoneal inoculations with the yeast phase of H. capsulatum, B. dermatitidis, and Candida (Monilia) albicans. With the dosages and strains of these fungi employed, it has been shown that:

1. Only 3 of 80 guinea pigs inoculated in this manner with large doses of the yeast phase of Histoplasma and 17 of 82 similarly inoculated with Blastomyces developed a generalized infection. Of 35 similarly inoculated with C. albicans, none developed such an infection.

2. There was no correlation between the size of the inoculating dose and the number of animals which developed generalized infection.

3. Relatively small doses of these fungi are as effective as larger ones in producing skin sensitization of these animals.

4. Both H. capsulatum and B. dermatitidis could be readily isolated from the spleens of apparently healthy guinea pigs which had been inoculated with variable doses of these fungi 9 to 10 months previously, even though there were no gross lesions apparent at autopsy. C. albicans could not be recovered from guinea pigs inoculated with this fungus 2 to 3 months previously.

REFERENCES

- DeMonbreun, W. A.: The cultivation and cultural characteristics of Darling's Histoplasma capsulatum. Am. J. Trop. Med. 14: 93 (1934).
 Hansmann, G. H., and Schencken, J. R.: A unique infection in man caused by a new yeast-like organism, a pathogenic member of the genus Sepedonium. Am. J. Path. 10: 731 (1934).
 DeMonbreun, W. A.: The dog as a patural host for Histoplasma cansulatum.

- donium. Am. J. Path. 10: 731 (1934).
 De Monbreun, W. A.: The dog as a natural host for Histoplasma capsulatum. Am. J. Trop. Med. 19: 565 (1939).
 Van Pernis, P. A., Benson, M. E., and Hollinger, P. H.: Specific cutaneous reactions with histoplasmin. J. A. M. A. 117: 436 (1941).
 Reid, J. D., Scherer, J. H., Herbut, P. A., and Irving, H.: Systemic histoplasmosis. J. Lab. & Clin. Med. 27: 419 (1942).
 Tager, M., and Liebow, A. A.: Observations on histoplasmosis. Induced infection in the mouse. Yale J. Biol. & Med. 14: 469 (1942).
 Henderson, R. G., Pinkerton, H., and Moore, L. T.: Histoplasma capsulatum as a cause of chronic ulcerative enteritis. J. A. M. A. 118: 885 (1942).
 Parsons, R. J., and Zarafonetis, C. J. D.: Histoplasmosis in man. Arch.
- rsons, R. J., and Zarafonetis, C. J. D.: Histoplasmosis in man. Int. Med. 75: 1 (1945). (8) Parsons,
- (9) Ricketts, H. T.: Oidiomycosis (blastomycosis) of the skin and its fungi.
- J. Med. Res. 6: 377 (1901).
 Bowen, J. T., and Wolbach, S. B.: A case of blastomycosis: the results of culture and inoculation experiments. J. Med. Res. 15: 167 (1906).
- (11) Davis, B. F.: The immunological reactions of oidiomycosis (blastomycosis) in the guinea pig. J. Inf. Dis. 8: 190 (1911).
 (12) Stober, A. M.: Systemic blastomycosis. Arch. Int. Med. 13: 509 (1914).
 (13) Long, E. R.: Specific necrosis in the infectious granulomas. J. A. M. A. 87:
- 1441 (1926).

- (14) Michelson, I. D., and Dulaney, A. D.: Experimental blastomycosis. South.
- (14) Michelson, I. D., and Dulancy, A. D..
 M. J. 24: 1034 (1931).
 (15) Benham, R. W.: The fungi of blastomycosis and coccidioidal granuloma. Arch. Dermat. & Syph. 30: 385 (1934).
 (16) DeMonbreun, W. A.: Experimental chronic cutaneous blastomycosis in monkeys. Arch. Dermat. & Syph. 31: 831 (1935).
- monkeys. Arch. Dermat. & Syph. 31: 831 (1935).

 (17) Ashford, B. K.: Further experimentation in animals with a Monilia com-
- (17) Ashford, B. K.: Further experimentation in animals with a Monital commonly found in sprue. Am. J. M. Sc. 131: 520 (1916).
 (18) Smith, L. W.: The role of M. psilosis (Ashfordi) in experimental sprue. J. A. M. A. 83: 1549 (1924).
 (19) Nye, R. N., Zerfas, L. G., and Cornwell, M. A.: The pathogenicity of yeast-like fungi isolated from the human gastro-intestinal tract. Am. J. M. Sc. 178: 515 (1929).
 (20) Repharm B. W.: Cortein propilies paresitie on man. J. Inf. Dis. 49:
- (20) Benham, R. W.: Certain monilias parasitic on man. 183 (1931).
- (21) Ninni, C., and Fittipaldi, C.: Quoted in: Owen, C. R., Anderson, M. B., and Henrici, A. T.: Allergy in *Monilia* and yeast infections. Mycopath. 8: 10 (1937)
- (22) Howell, A.: The efficiency of methods for the isolation of Histoplasma capsulatum. Pub. Health Rep. 63: 173 (1948).
 (23) Howell, A.: Studies of fungus antigens. I. Quantitative studies of cross reactions between histoplasmin and blastomycin in guinea pigs. Pub. Health Rep. 62: 631 (1947).
- (24) Conant, N. F.: A cultural study of the life-cycle of Histoplasma capsulatum.
- J. Bact. 41: 563 (1941).
 (25) Martin, D. S.: Studies on the immunologic relationships among various species of the genus Candida (Monilia). Am. J. Trop. Med. 22: 295
- (26) Emmons, C. W., Bell, J. A., and Olson, B. J.: Naturally occurring histoplasmosis in Mus musculus and Rattus norwegicus. Pub. Health Rep. 62: 1642 (1947).

UNITED NATIONS WORLD HEALTH ORGANIZATION INTERIM COMMISSION

NATIONS UNIES ORGANISATION MONDIALE DE LA SANTE COMMISSION INTÉRIMAIRE

EXPERT COMMITTEE ON TUBERCULOSIS

Report of the First Session

Office International d'Hygiène Publique, Paris.

July 30-August 2, 1947

At the third session of the Interim Commission of the World Health Organization held in Geneva in April 1947, it was resolved to set up an Expert Committee on Tuberculosis. The chairman of the Interim Commission and the executive secretary agreed to appoint the following members of this committee, after approaching their respective Governments:

Dr. P. D'Arcy Hart, Medical Research Council, London.

Dr. Herman E. Hilleboe, United States Public Health Service,1 Washington, D. C.

Appointed Commissioner of Health, New York State, July 1, 1947.

Dr. Johannes Holm, State Serum Institute, Copenhagen.

An invitation to the Government of the U. S. S. R. to suggest the name of a Russian member of the committee was sent. The attendance of an expert was arranged, but his sudden illness prevented his coming to Paris.

The Expert Committee on Tuberculosis appointed by the Interim Commission met in Paris from July 30 to August 2, 1947.

Dr. Holm was elected chairman. The Secretary to the Expert Committee, Dr. J. B. McDougall, was in attendance.

The following report of the Committee was accepted by the Interim Commission at its fifth session for submission to the World Health Assembly:

A. Introduction

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It is recognized that tuberculosis is a world problem of great magnitude. The committee is fully in accord with the decision of the Interim Commission that tuberculosis, malaria, and venereal disease are infectious diseases deserving the highest priorities for its activities.

There can be no isolationism in the field of health. The fight against infectious disease is not a national or a racial problem; it is a task for the whole of humanity. No nation is safe if another nation is vanquished by disease. The fortunate and relatively healthy nations, inspired by intelligent self-interest and humane considerations, will necessarily have to come to the aid of stricken nations, and through money, professional personnel and equipment, distribute existing resources to the needy and suffering areas of the world.

Tuberculosis control work of an international scope must go forward if present suffering and disability are to be alleviated and future generations protected. The all-inclusive objective of any sound tuberculosis program is the prevention and eventual eradication of tuberculosis from the peoples of the world. Poverty, shortages of food and housing, and the lack of opportunity for gainful occupation, complicate the task enormously, and make it necessary for us to share and distribute our resources where they will do the most good in the shortest possible time.

B. Fields of Activity

There are five well-defined fields of activity in which we must work and direct our efforts on a planned basis if tuberculosis is to be systematically eliminated: (1) Prevention, (2) case finding, (3) isolation and medical care, (4) rehabilitation and after-care, (5) social and economic protection of afflicted families.

No one of these activities can be effective alone. They all must operate together and in proper sequence.

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C. Techniques for Control

It is not enough merely to recognize and describe the objectives of a tuberculosis control program. It is also necessary to have clearly defined and firmly established techniques for the achievement of those objectives. The following recommendations include 11 principal techniques for tuberculosis control, which may be used singly, in groups, and finally all together, if the World Health Organization

program is to be comprehensive and wholly effective:

1. The first technique is the determination of the extent of the problem of tuberculosis in each country, the present means and facilities of its disposal, the manner in which these facilities are being used to tackle the problem, and the additional facilities needed. Countries with little information available should be encouraged to record at least simple basic data. It is recommended that schedules (now being prepared by the Expert Committee) be filled in by the experts of the Secretariat, who, on request, actually go into the countries. These schedules should be kept up to date at regular intervals.

2. One of the most important techniques that works toward the realization of the objectives of tuberculosis control is the recruitment and training of professional personnel. In most countries there is at present an insufficient number of well-trained workers in this field. It is recommended that traveling fellowships be awarded to countries most in need, principally to train medical officers. are four special fields in which trained medical officers are essential for every country-administration, epidemiology, laboratory work, and clinical work. It is estimated that one thousand such fellowships could be granted by the World Health Organization with good effect within the next few years. It would appear wise to recommend that only 50 of these be provided in the first year in order to get the program under way. To operate the scheme, the Secretariat should survey the teaching facilities throughout the world and designate acceptable teaching centers. At the same time, the Secretariat should ascertain the needs of countries for trained personnel, so that after such consultation, promising medical officers, especially those who show potentialities of leadership, can be selected for fellowships. Countries where the needs are greatest should be chosen first.

It is also recommended that consultation services of short duration be provided to countries, especially those with teaching centers, in order to make available the latest knowledge and viewpoints of out-

standing specialists.

3. The provision of physical facilities, supplies and equipment for all phases of prevention, diagnosis and treatment is second in importance only to the provision for personnel. It is recommended that the World Health Organization should be prepared to give expert advice to the various countries requesting such information, on the

number, type and location of facilities needed, and on the best means of financing the construction and maintenance of these facilities, drawing on the successful experience of other countries. Recommendations should be given only if they are to form a part of a long-range, comprehensive plan for the nation and its administrative subdivisions.

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4. Heath education is recognized as an essential tool in tuberculosis control. The general public must know the seriousness of the disease and its cost in human misery and money before it will accept its responsibility to support the work financially. It is recommended that the World Health Organization should encourage national and international voluntary organizations to take the major responsibilities for informing the public and gaining their support.

To keep the medical profession informed on advances in tuberculosis, it is recommended that the World Health Organization prepare from time to time material on recent developments of special importance, and that the circulation of specialist literature be provided. The World Health Organization should encourage national and international professional organizations to develop the distribution of tuberculosis literature.

5. The best way to get a new program started, or to improve a poor one, in any country is by means of field services for the purpose of demonstrating practical activities in one or more of the special fields of administration, epidemiology, laboratory work and clinical Well-trained teams, even with limited supplies and equipment, can demonstrate what should be done to control tuberculosis and how It is recommended that the World Health Organization provide demonstration teams. The size of the team and the length of its stay would vary with needs, but in any event should be kept to a minimum. Certain supplies and equipment will be necessary for An essential condition for the demonstration will be that the country agrees beforehand to take over the project as soon as sufficient of its personnel has been trained to do so. When taken over, these field demonstrations should become national training centers, and in some cases should be designated also for international use of traveling fellows. For example, an international training center might be established in India for training workers from various parts of Asia, where the problems to be solved are similar in nature. Areas where it is proposed to set up international training centers should have first call on demonstrations, if such are necessary.

The persons charged with these demonstrations could be either regular staff members of the World Health Organization or professional personnel with temporary appointments. The person to take charge of the work, when it is taken over by the local group, could well be one of the persons who had received a traveling fellowship from the World Health Organization.

- 6. While it is recognized that present budgetary limitations do not permit grants of money for tuberculosis control to nations at this time, it is recommended that in future such grants should be made, in order to help nations unable to help themselves. Such grants should be made, however, only if great need is demonstrated, a complete plan submitted to show the joint use of national funds and those from the World Health Organization, and to show that the funds are used solely for tuberculosis control and that the World Health Organization's contributions are not used to replace local funds.
- 7. The best contribution that can be made by the World Health Organization in tuberculosis research would appear to be in developing and recommending uniform procedures. Special problems would require from time to time the services of small subcommittees of experts in highly specialized fields. Where possible, members of other expert committees of the World Health Organization should be used for this purpose. Whenever a problem comes up for the Expert Committee on Tuberculosis which involves the responsibility shared by another expert committee, one of the members of the second committee should be asked to take part in the delibera-For example, when the Expert Committee on Tuberculosis considers the problem of tuberculin and tuberculin testing a member of the Expert Committee on Biological Standardization should be asked to participate, and vice versa. It is recommended that the Committee on International Classification of Morbidity and Mortality consult with the Expert Committee on Tuberculosis before final action is taken on classification of tuberculosis. several suggestions our Committee wishes to make on the first draft.

The principal problems which need action to establish uniform procedures are as follows:

- (1) Tuberculin and tuberculin testing.
- (2) Preparation and clinical use of BCG.
- (3) Classification of tuberculosis.
- (4) X-ray interpretation and mass radiography.
- (5) Laboratory diagnosis of tubercle bacilli.
- (6) Evaluation of new chemotherapeutic agents such as streptomycin.

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Even during the period of the Interim Commission it is recommended that action be taken on (1), (2), and (6). Thus, it is urged that subcommittees be appointed on tuberculin and tuberculintesting, and on BCG, and that a conference be called early in 1948 on the use and value of streptomycin. This conference should bring together those who have been actively engaged in research on this drug.

8. It is recognized that several other international organizations have been carrying on activities and have contributed in many ways to tuberculosis control. It is recommended that the World Health Organization should take full advantage of these services and should establish working relationships with all groups genuinely interested in tuberculosis control. Such a cooperative effort would help to avoid duplication and would produce harmonious agreement in this collective enterprise. The Committee has been informed that the International Union Against Tuberculosis is about to establish a branch office in Geneva. It is urged that liaison be established at once between World Health Organization and the Union in order that their several activities go forward in unison. Cooperation with all private and official agencies, even those only partially engaged in tuberculosis control activities, should be extended at every oppor-Furthermore, this Committee would welcome the opportunity to be consulted by other committees of the World Health Organization and United Nations whenever questions and problems involving tuberculosis arise.

9. Tuberculous cattle still form an important source of tuberculosis among human beings throughout the world. Infected milk is not the only source of spread, for it has recently been demonstrated that farm workers may contract bovine tuberculosis through direct contact with diseased cattle. It is recommended that the World Health Organization use its influence to encourage nations whose herds have high infection rates to take active steps to eradicate

tuberculosis among cattle as quickly as possible.

10. It is recommended that the World Health Organization be prepared to give expert advice to national governments and health departments on sound laws and regulations pertaining to human and bovine tuberculosis. This committee proposes to study both the legal and epidemiological aspects of the problem of tuberculosis among migrants. This would form the basis of recommendations designed to prevent the spread of this disease from one country to another.

11. Modern public health practice demands that public health programs have review and evaluation at regular intervals, in order that any ineffective techniques may be discarded and that more modern ones be added as new knowledge is gained. This is particularly true of a new program. Accordingly, it is recommended that the World Health Organization make preparation for review and evaluation of its program at yearly intervals with the advice and counsel of the Expert Committee.

D. Emergency Measures

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Because of the epidemic proportions of tuberculosis in many

May 7, 1948 622

countries, certain emergency measures, which require relatively small expenditures, should be applied at once. It is recommended that small demonstration teams be sent into such countries, even for short periods, to carry on intensive programs of BCG vaccination similar to those at present in operation under the Danish Red Cross in several European countries which have appealed for aid.

The Committee wishes to emphasize that this measure is clearly of an emergency nature. It is hoped that the initiation and successful operation will encourage the local groups to develop and carry on a

more comprehensive program.

E. Tuberculosis Secretariat and Finance

In order to accomplish the above proposals, it is recommended that a permanent Tuberculosis Control Office be established within the World Health Organization.

Even though no funds are available yet for all these proposals, which the Committee hopes the World Health Organization will in due course accept, it is recommended that some funds be provided immediately by the Interim Commission for the emergency measure under heading D, namely, to start in certain countries, as soon as possible, programs for BCG vaccination and for identifying infectious cases.

It is further recommended that the Interim Commission provide immediate funds for the expenses of the subcommittee meetings (on tuberculin and tuberculin testing, and on BCG) and for the conference on streptomycin, referred to under heading C7.

F. Dissemination of Information

If the World Health Organization Interim Commission approves of the proposals of this Committee, it is recommended that there should be wide dissemination of information concerning the services the World Health Organization can provide.

INCIDENCE OF COMMUNICABLE DISEASES IN THE UNITED STATES February 29-March 27, 1948

The accompanying table summarizes the incidence of nine important communicable diseases, based on weekly telegraphic reports from State health departments. The reports from each State for each week are published in Public Health Reports under the section "Incidence of Disease." The table gives the number of cases of these diseases for the 4 weeks ended March 27, 1948, the number reported for the corresponding period in 1947, and the median number for the years 1943–47.

DISEASES ABOVE MEDIAN INCIDENCE

Influenza.—The number of cases of influenza dropped from 45,536 during the preceding 4 weeks to 25,459 during the 4 weeks ended March 27. Of the total cases, Texas reported 11,620, South Carolina 2,825, Virginia 2,213, Alabama 1,512, California, 1,447, Arkansas 985, and Arizona 834; 84 percent of the cases occurred in those 7 States. For the country as a whole the incidence was only about 20 percent of the 1947 incidence for the corresponding weeks, but it was 1.4 times the median for the preceding 5 years. The recent outbreak of this disease that has been confined to States in the Southern and Western sections followed the usual pattern of an influenza epidemic and the peak apparently was reached during the month of February. The 1947 epidemic did not begin until late in the season and the peak was not reached until in March, or during the 4 weeks corresponding to the period under consideration.

DISEASES BELOW MEDIAN INCIDENCE

Diphtheria.—The incidence of diphtheria continued to decline. For the 4 weeks ended March 27 there were 732 cases reported as compared with a median of 1,067 cases for the corresponding period in the preceding 5 years (1943–47). In the South Atlantic and Mountain sections the incidence was somewhat above normal, but in all other sections the numbers of cases were relatively low. For the country as a whole the current incidence was the lowest in the 20 years for which data are available in this form.

Measles.—For the current 4-week period there were 83,160 cases of measles reported. The number was more than 3 times that reported for the corresponding period in 1947, but it was slightly below the median for the preceding 5 years. The disease was most prevalent in the East North Central and West South Central sections; in the former section the number of cases (29,760) was 2.1 times the 1943–47 median and in the latter section the number of cases (8,364) was 1.5 times the normal seasonal median. A slight increase was reported from the Pacific section but in the other 6 sections the incidence was considerably below the expected seasonal incidence.

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Meningococcus meningitis.—The incidence of meningococcus meningitis (352 cases) was the lowest reported for this period since 1942 when 329 cases were reported for the corresponding 4 weeks. The New England, South Atlantic, and Pacific sections reported more cases than occurred during the same weeks in 1947, but the incidence in all sections was considerably below the median for the preceding 5 years. However, since the 1943-47 median fell within an epidemic period of this disease a better comparison is with the average incidence for nonepidemic years which is approximately 300 cases.

Poliomyelitis.—The number of cases (110) of poliomyelitis reported during the 4 weeks ended March 27 compared very favorably with the median for the preceding 5 years (112 cases). The median for this disease also falls within an epidemic period and, while the current incidence is about the same as the median, the number of cases is still higher than the average of 80 cases for nonepidemic years. While the numbers of cases were not large in the West North Central and Mountain sections (16 and 15, respectively) they were more than twice the medians for those sections. Minor increases were reported from 3 other sections and in the other 4 sections the incidence was lower than the median.

Scarlet fever.—The incidence of scarlet fever continued at a relatively low level, the current incidence (10,546 cases) being about 85 percent of the 1947 incidence for the corresponding period and 61 percent of the median for the preceding 5 years. While each section of the country has shared in the favorable situation for this disease that now exists, the greatest declines from the normal incidence during this particular 4-week period occurred in the New England, South Atlantic, and Mountain sections. Since 1944 this disease has been on the downward swing of a long-term cycle and for the current 4-week period the incidence was the lowest on record.

Smallpox.—Eight cases of smallpox were reported during the 4 weeks ended March 27, as compared with 19 in 1947 and a median for the preceding 5 years of 39 cases. Three cases occurred in Kansas and one each in Minnesota, North Dakota, North Carolina, Georgia, and Oklahoma. The number of cases was the lowest on record for these same weeks.

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Typhoid and paratyphoid fever.—For these diseases the incidence (166 cases) during the 4 weeks ended March 27 was the lowest for a corresponding period in the 20 years for which these data are available. A few more cases than might be expected occurred in the West North Central and West South Central sections, but in other sections the incidence was about equal to or lower than the seasonal expectancy. The greatest declines were reported from the North Atlantic and East North Central sections.

Whooping cough.—The incidence of this disease was also relatively low, the 9,485 cases reported being less than 90 percent of the median

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for the preceding 5 years. Of the 9 geographic sections, 4 reported an increase over the normal seasonal median and in 5 sections the incidence was lower than the expectancy. The greatest increase in cases was reported from the West South Central sections and the greatest decline occurred in the Middle Atlantic section.

MORTALITY, ALL CAUSES

For the 4 weeks ended March 27 there were 39,148 deaths from all causes reported to the National Office of Vital Statistics by 93 large cities. The median number of deaths for the corresponding period in the years 1945–47 was 38,603. The number of deaths was higher than the preceding 3-year median in each week of the current 4 weeks except the first one and for the 4 weeks the number was 1.4 percent above the median.

Number of reported cases of 9 communicable diseases in the United States during the 4-week period February 29-March 27, 1948, the number for the corresponding period in 1947, and the median number of cases reported for the corresponding period, 1943-47

Division	Cur- rent period	1947	5-year median	Cur- rent period	1947	5-year median	Cur- rent period	1947	5-year median
	D	iphther	ia	1	Influenza	1		Measles	
United States	732	1,068	1,067	25, 459	125, 077	17,615	83, 160	27, 064	87, 789
New England	24	85	40	37	52	83	3, 773	6, 787	6, 787
Middle Atlantic	77	131	131	166	90	90	16, 981	4,608	21, 783
East North Central	99	145	142	312	2, 620	710	29, 760	4, 568	13, 993
								639	7, 371
West North Central	76	111	111	246	17,063	183	5, 648		7,371
South Atlantic	152	138	138	5, 574	15, 939	4, 540	4, 488	4, 526	5, 222
East South Central	67	121	108	2, 111	3, 933	1,391	2,098	1, 195	3, 863
West South Central	88	164	204	13, 462	76, 571	6, 921	8, 364	2, 161	5, 634
Mountain	92	55	52	1,472	7, 751	1,144	3, 397	1,398	4, 969
Pacific	57	118	118	2,079	1,058	553	8, 651	1, 182	8, 469
		ningocoo		Pe	oliomyeli	tis	Sc	earlet fev	er
United States	352	372	1,018	110	156	112	10, 546	12, 272	16, 287
New England	18	11	45	1	6	6	924	879	2, 361
Middle Atlantic	58	70	239	14	12	12	2, 964	3, 112	4, 844
East North Central	51			12	18	9	3, 480	4, 085	4, 413
		52	199			7		1.082	
West North Central	22	42	70	16	16		834 671		1,718
South Atlantic	52	44	158	13	14	14		818	1, 522
East South Central	39	39	93	9	12	8	289	553	553
West South Central	51	64	101	11	22	17	274	313	465
Mountain	7	8	13	15	7	7	326	498	848
Pacific	54	42	111	19	49	23	784	932	1,054
	8	mallpox			phoid ar		Who	oping co	ugh
United States	8	19	39	166	189	198	9, 485	10, 709	10, 667
New England	0	19		6	15	13	780	1,002	1, 168
Middle Atlantic	0	0	0	21		33	1.369	1, 984	1, 984
Fact North Control			0		20				1, 569
East North Central	0	5	6	15	29	29	1,661	2, 451	
West North Central	5	8	9	14	10	10	464	345	377
South Atlantic	2	1	2	37	30	38	1, 232	1,384	1, 521
East South Central	0	1	5	15	15	16	404	427	435
West South Central	1	2	11	37	30	34	2, 172	2,006	1, 196
Mountain	0	2	2	8	8	9	645	237	399
Pacific	0	0	2	13	32	20	758	873	873

¹ New York, North Carolina, and Pennsylvania excluded; New York City and Philadelphia included.

INCIDENCE OF DISEASE

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No health department, State or local, can effectively prevent or control disease without knowledge of when, where, and under what conditions cases are occurring

UNITED STATES

REPORTS FROM STATES FOR WEEK ENDED APRIL 17, 1948 Summary

A total of 32 cases of poliomyelitis (the same number as for the corresponding week last year) was reported for the current week, as compared with 37 last week and a 5-year (1943-47) median of 28. Only 2 States reported more than 2 cases—Texas 6 (last week 9), and Michigan 3 (last week 2). The total reported for the 4 weeks since the average date of seasonal low incidence is 120 cases, as compared with 124 for the same period in 1945, the highest number reported for a corresponding period of the past 5 years, 68 in 1944 the lowest, and a 5-year median for the period of 108 cases, reported in 1946.

For the current week, 25,616 cases of measles was reported, as compared with 25,842 last week (the highest incidence so far this year) and a 5-year median of 27,161. In 1946 the peak of incidence was reached in the week ended April 13 with a reported total of 40,746 cases. The total for the year to date is 250,733, as compared with 82,918 for the same period last year and 262,946 for the 5-year median.

Of the total of 2,044 reported cases of influenza (last week 2,702, corresponding week last year 23,536, 5-year median 1,917), 1,514 cases were reported in 4 States—Texas 879, South Carolina 283, Virginia 246, and Oklahoma 106.

Four cases of Rocky Mountain spotted fever were reported, 1 each in New Jersey, Wyoming, Colorado, and Oregon. New York reported 2 cases of anthrax, and New Jersey 1 case. One case of smallpox was reported, in Nebraska, and 1 case of psittacosis, in Michigan.

During the week 8,942 deaths from all causes were recorded in 92 large cities in the United States, as compared with 9,601 last week, 9,670 and 9,051 respectively, for the corresponding weeks of 1947 and 1946, and a 3-year (1945–47) median of 9,078. The total for the year to date is 160,698, as compared with 160,962 for the corresponding period last year. Infant deaths in the same cities totaled 655, as compared with 707 last week and a 3-year median of 631. The cumulative total is 11,006, as compared with 12,738 for the same period in 1947.

Telegraphic morbidity reports from State health officers for the week ended Apr. 17, 1948, and comparison with corresponding week of 1947 and 5-year median

In these tables a zero indicates a definite report, while leaders imply that, although none was reported, cases may have occurred.

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	D	iphthe	ria	1	nfluenz	8		Measles	1		eningi ningoco	
Division and State	We	eek ed—	Me-	We	eek ed—	Me-	We		Me- dian	wende	eek ed—	Me-
	Apr. 17, 1948	Apr. 12, 1947	dian 1943- 47	Apr. 17. 1948	Apr. 12, 1947	dian 1943- 47	Apr. 17, 1948	Apr. 12, 1947	1943- 47	Apr. 17, 1948	Apr. 12, 1947	1943-
NEW ENGLAND												
Maine	0	0	0		4	2	6	250	91	0	0	
New Hampshire	0	0	0		8 15	0	17	9 285	9 156	0	0	
Vermont	0	9	8		10	0	1, 373	360	1,013	2	ĭ	
Rhode Island	ő	1	ő			1	9	206	14	0	0	
Connecticut	3	0	0	3	9	3	146	864	430	0	1	
MIDDLE ATLANTIC												
New York	2	12		16	1 18	12	2, 252	532	2, 317	6		17
New Jersey	3 2	3 17	6 12	(1) 3	16 2 4	12	1, 165 1, 587	451 283	1, 831 898	1 3	1 7	13
Pennsylvania	-	1,	12	(-)	- 4		1,001	200	000		1	-
EAST NORTH CENTRAL	5	10	13	1	32	14	1,098	582	916	3	13	13
OhioIndiana	10	4	3		17	7	668	120	224	ő	0	1
Illinois	5	4	15	2	16	16	1, 805	92	1, 281	2	16	10
Michigan 3	1	4	8	<u>i</u>	11	1	1, 479	69	812	2	7 2	1
Wisconsin	1	4	3	1	290	36	1, 891	326	2, 277	1	2	1
WEST NORTH CENTRAL							400	00	00			
Minnesota	0	0	2 4		1,576	2	462 359	96 181	96 181	0	4 2	1
Iowa	2	25	4	17	7,575	3	550	6	73	3	3	1
North Dakota	0	0	ō		2	2	16	7	8	0	1	(
South Dakota	0	1	1	7		i	45	12	38	0	0	0
Nebraska	1	2	1 3	7	72	5	244 88	8	166 623	0 2	1 0	1
Kansas	1	4	9	******	12		00	-	020	-		
SOUTH ATLANTIC Delaware	0	0	0				42		12	0	1	,
Maryland 3	7	14	6	5	11	6	145	23	168	1	5	10
District of Columbia.	Ö	0	0			1	169	44	83	0	2	2
Virginia	1	4	4	246	4, 673	274	145	288	488	4	3	8
West Virginia North Carolina	6	6	6	11	935	12	324 16	22 221	116 221	0	3 5	
South Carolina		9	5	283	2, 650	292	77	210	251	2 2	ő	10 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2
Georgia	2	5	1	19	485	43	39	155	216	0	0	4
Fiorida	5	2	3	35	109	16	269	127	127	0	1	
EAST SOUTH CENTRAL												
Kentucky	2	8	5	5		4	179	3	81	3	3	7
Tennessee	4 2	5	5	44 95	741 727	43 87	212 61	96 188	286 235	0	3 5	2 2 2
Mississippi 3	3	3	5	16	118		56	16		1	1	2
WEST SOUTH CENTRAL												
Arkansas	0	3	2	73	1, 255	35	82	113	193	1	2	2
Louisiana	3	2 3	2 3	5	300	16	74	26	84	1	0	2
Oklahoma	12	18	29	106 879	3, 347 3, 896	79 778	2, 927	11 374	1, 297	1	10	10
Texas MOUNTAIN	12	19	28	919	3, 000	****	2, 021	013	1, 201		10	10
	3	0	1	2	571	4	28	162	124	0	0	0
MontanaIdaho	0	0	ó	21	63	2	121	12	67	ő	0	i
Wyoming	0	0	Ö	1			116	19	60	0	0	1
Colorado	3	9	4	22	641	25	641	93	225	1	0	0
New Mexico	4	8	0	8 36	165	78	35 224	77 81	24 81	0	0	1
Arizona Utah ³	2	1	0	1	98	3	205	13	164	0	0	1
Nevada	0	0	0						11	0	0	0
PACIFIC												
Washington	0	2	5	4	480	3	761	28	133	1	2	3
Oregon	0	1	7	34	85	13	4 87	27	1 526	0	0	22
California	7	20	20	53	79	29	3, 258	178	1, 536	3	10	_
Total	116	230	244		23, 536	1, 917	25, 616		27, 161	49	122	194
15 weeks	2,976	4, 231	4, 231	126, 054	266, 137	179, 321	250,733	82, 918	262, 946	1, 249	1, 322	3, 617
Seasonal low week	(27th) July	5-11	(30th) J	uly 26-	Aug. 1	(35th) A	ug. 30-	Sept. 5	(37th)	Sept.	13-19

New York City only.
 Philadelphia only.
 Period ended earlier than Saturday.
 Observed in Arizona for week ended April 3—included in cumulative totals only.
 Dates between which the approximate low week ends. The specific date will vary from year to year.

Telegraphic morbidity reports from State health officers for the week ended Apr. 17, 1948, and comparison with corresponding week of 1947 and 5-year median—Continued

	Pol	iomyel	itis	Sca	rlet fev	er	S	mallpo	x	Typh	oid and hoid fe	para ver
Division and State	We	eek ed—	Me-	We	ek ed	Me-	Wende	eek ed—	Me- dian	We		Me- dian
	Apr. 17, 1948	Apr. 12, 1947	dian 1943- 47	Apr. 17, 1948	Apr. 12, 1947	dian 1943- 47	Apr. 17, 1948	Apr. 12, 1947	1943- 47	Apr. 17, 1948 7	Apr. 12, 1947	1943-
NEW ENGLAND		_										
Maine	0	1	0	8 5	21	34	0	0	0	4	0	9
New Hampshire	0	0	0	0 5	10	10	0	0	0	0	0	9
Vermont	0	0	0	182	115	366	0	0	ő	5	6	
Rhode Island Connecticut	0	0	0	5 18	16 42	22 71	0	0	0	1 0	0	1
MIDDLE ATLANTIC												
New York	2	7	4	6 269	343	535	0	7	0	1	0	
New Jersey	1	2	1	67 308	155 216	155 405	0	0	0	0 3	0	
Pennsylvania	0	0	1	308	210	400	U	0	0			1
EAST NORTH CENTRAL				211	950	950	0	0	1	4	2	,
OhioIndiana	1	0	0	311	358 71	358 105	0	1	1	1	2	
Illinois	0	1	1	107	103	203	0	0	1	1	0	
Michigan	3	1	1	123	128	160	0	0	0	2	2	
Wisconsin	0	0	0	69	39	168	0	0	0	0	1	(
WEST NORTH CENTRAL												
Minnesota	2	0	0	29	48	71	0	0	0	0	0	(
Missonsi	0 2	0	0	8 33	40 38	57 56	0	0	0	1	o	(
Missouri North Dakota	0	0	0	3	7	8	0	o	0	1	0	(
South Dakota	0	Ö	ő	1	10	19	0	0	0	0	0	(
Nebraska	2	0	0	17	13	38	1	0	0	0	0	(
Kansas	0	0	0	25	43	72	0	0	0	1	0	(
SOUTH ATLANTIC												
Delaware	0	0	0	3	7	7	0	0	0	0	0	(
Maryland	0	0	0	6 34 8	28 8	148 24	0	0	0	3	0	(
District of Columbia Virginia	0	0	0	21	32	82	0	ő	ő		o	1
West Virginia	ő	0	0	13	11	39	0	0	0	8	0	(
North Carolina	2	0	0	19	20	39	0	0	0	1	0	(
South Carolina	0	0	0	17	2	3	0	0	0	0 2	0	
Georgia	2	0 2	0	6	14 13	14 10	0	0	0	2	i	1
Florida		-	-		10	10						
	0	3	0	10	26	38	0	0	0	1	2	- 1
Kentucky Tennessee	0	1	ő	20	42	38	ő	ő	Õ	2	0	1
Alabama	ő	o	ĭ	6	6	16	0	0	0	1	0	(
Mississippi *	0	1	1	2	4	10	0	1	0	1	0	(
WEST SOUTH CENTRAL												
Arkansas	0	0	0	4	5	6	0	0	0	2 2	0	(
Louisiana	0	0	0	9	7	7 16	0	0	0	ő	0	i
Oklahoma Texas	6	ő	3	29	31	63	0	0	1	5	1	1
MOUNTAIN					-							
Montana	0	0	0	22	3	15	0	0	0	0	0	(
Idaho	1	0	Ö	6 14	7	28	0	0	0	0	0	
W yoming	0	0	0	2	2	22	0	0	0	0	0	(
Colorado	0	0	0	15	43 11	45	0	0	0	0	0	1
New Mexico	0	0	0	4 6	11	11 13	0	0	0	0	0	1
Utah 3	0	0	ő	16	15	30	0	0	0	0	0	(
Nevada	0	0	0	0	0	0	0	0	0	0	0	(
PACIFIC											_	
Washington	2	1	1	53	28	44	0	0	0	0	0	1
Oregon	1 2	0	0 5	12 104	129	41 201	0	0	0	0	2	
California		10	-					- 9	10	58	32	58
Total	32	32	28	2, 088	2, 381	4, 483	37			689	641	802
	468	721	521	34, 969	40, 604	59, 767	_	77				
Seasonal low week 5	(11th) Mar.	15-21	(32nd	l) Aug.	9-15	(35t)	h) Aug Sept. 5	. 30-	(11th)	Mar.	15-21
Total since low	120	109	108	57, 508	67, 290	98, 088	58	131	243	216	156	210

Period ended earlier than Saturday.
 Dates between which the approximate low week ends. The specific date will vary from year to year.
 Including cases reported as streptococcal sore throat.
 Including paratyphoid fever reported separately, as follows: Massachusetts 4 (salmonella infection), Virginia 1, Tennessee 1, California 1.

Telegraphic morbidity reports from State health officers for the week ended Apr. 17, 1948, and comparison with corresponding week of 1947 and 5-year median—Continued

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on),

	Who	ooping	cough			Wee	k ende	d Apr. 1	7, 1948		-
District and Otto	Week	ended-	Me-	1	ysent	ery	En-	Rocky	1	Ty-	Un-
Division and State	Apr. 17, 1948	Apr. 12, 1947	dian 1943- 47	Ame	Bacil lary		ceph- alitis, infec- tious	spot-	Tula- remia		du- lant
NEW ENGLAND											
Maine New Hampshire	11 1 59	15			30						
Vermont	47	124			1		1				
Rhode IslandConnecticut	1 25	36	18	3			1				
New York	101	115		16							
New Jersey Pennsylvania	64 62	129 163					*****	1			
EAST NORTH CENTRAL										*****	
Ohio	74	144								*****	1
IndianaIllinois	35 40	48 54			6		*****				1
Michigan 3	85	148		14							4
Wisconsin	110	127	104						1		11
Minnesota	26	23	10	2							
Iowa	11	10	11								2
Missouri	30	20	11					*******			1
North Dakota	9 7	1	2	1						*****	
Nebraska	3	6	6							*****	
Kansas	45	18	28						1		10
Delaware		3	1								
Maryland 3	20 7	58	58			2				*****	9
District of Columbia	7	7	7	1							
Virginia.	36 16	84	84			48					
West Virginia North Carolina	99	10 13	26 94					******	1	9	
South Carolina	81	108	79	1	3		1			1	1
GeorgiaFlorida	18 28	30 48	30 24	1	1				3		3
EAST SOUTH CENTRAL											
Kentucky	7	9	34								
Tennessee	42	25	29	1		1					1
Alabama	40	38	18	1					*****	1	
Mississippi 3		5		1	1				*****		
Arkansas	23	17	11	16		2			2		2
Louisiana	11	20	3	2					4		1
Oklahoma Texas	39 502	13 583	10 351	14	402	33				2	8
MOUNTAIN											
Montana	6	1	2								
daho	6	10	6					1			
Colorado	56	33	33					1	*****		7
New Mexico	21	21	7			2					
Arizona	68 18	12	18			6					1
Utah	10	6	33								
PACIFIC											
Washington	33	18	27	1		2					1
Oregon	23 97	10	18	9				1			3
California	97	141	141	6	4						3
Total	2, 149	2, 528	2, 528	100	461	96	3	4	12	6	107
Same week: 1947	2, 528			42	187	146	8	1	14	36	99
Median, 1943-47	2, 528			36	261	74	8	2	13	36	1 99
	32, 906			997	3, 977	2,852	129	10	266		1, 369
	38, 015 36, 627			697	4, 612 4, 252	3, 171	101	13	525 267	614	1, 564 1, 287

Period ended earlier than Saturday.
 9 cases undulant fever reported in February and March in error; deducted from cumulative totals.
 Alaska: Chickenpox 3, measles 1, influenza 2, mumps 1, pneumonia 2, whooping cough 25. Territory of Hawaii: Rabies 0, measles 5, scarlet fever 3, typhus fever (endemic) 2, whooping cough 44. Anthrax: New York 2, New Jersey 1. Leprosy: Florida 2. Psittacosis: Michigan 1.
 3-year median, 1945-47.

WEEKLY REPORTS FROM CITIES*

City reports for week ended Apr. 10, 1948

This table lists the reports from 89 cities of more than 10,000 population distributed throughout the United States, and represents a cross section of the current urban incidence of the diseases included in the table.

	cases	tis, in-	Influ	enza	8	me-	nía	litis	ever	898	and	cough
Division, State, and City	Diphtheria	Encephalitis, fectious, cas	Cases	Deaths	Measles cases	Meningitis, meningococcus, cases	Pneumon deaths	Poliomyelitis cases	Scarlet fev	Smallpox cases	Typhoid and paratyphoid lever cases	Whooping conses
NEW ENGLAND												
Maine: Portland	0	0		0	2	0	2	0	3	0	0	
New Hampshire: Concord	0	0		0	3	0	2	0	1	0	0	
Massachusetts: Boston	1	0		0	493	0	9	0	55	0	0	9
Fall River	0	0		0	5	0	0	0	1 3	0	0	3
Springfield	0	0		0	1	0	14	0	11	0	0	2
Rhode Island:	0	0	2	1	1	0	3	0	4	0	0	
Providence Connecticut:			-		1	0	0	0	2	0	0	
Bridgeport	0	0		0		0	0	0	2	0	0	
New Haven	0	0		0	2	ő	3	ő	2	0	0	3
MIDDLE ATLANTIC												
New York: Buffalo	0	0		0	13	1	1	0	7	0	0	5
New York	8	0	4	1	1, 544	5	79	0	56	0	0	21
Rochester	0	0		0	3	0	3	0	10	0	0	6
Syracuse New Jersey:	0	0	*****	0	3	0	3	0	5	U	0	
Camden	1	0		0	14	0	0	0	1	0	0	
Newark	0	0	1	1	167	0	0	0	5	0	0	
TrentonPennsylvania:	1	0	-*	0	4	1	4	0	1	U	0	•
Philadelphia	3	0	2	0	703	0	17	0	41	0	0	19
Pittsburgh	0	0	1	1	8	1	9	1	26	0	0	6
Reading	0	0		0	9	0	3	0	14	0	0	3
EAST NORTH CENTRAL												
Ohio:												
Cincinnati	1 0	0	1	1	87 16	2	6	0	38	0	0	3 13
Cleveland	0	0	1	0	57	Ô	2	0	6	ő	0	2
Indiana:	-											
Fort Wayne	0	0		0	23 179	0	2 2	0	7 2	0	0	12
Indianapolis South Bend	0	0		0	6	0	0	0	2	0	0	
Terre Haute	0	0		0		0	2	0	0	0	0	1
Ulinois:	1	0		0	977	3	24	0	48	0	0	11
Chicago	Ô	ő		0	12	0	3	0	1	Ö	0	
Michigan:		0			900	1		0	49	0	0	19
Detroit	2	0		0	298	0	8	0	49	0	0	
Grand Rapids	0	ő		0	57	Ö	2	0	i	0	0	2
Wisconsin:					m.,	0	0	0	0	0	0	
Kenosha	0	0		0	78 48	0	3	0	18	0	0	2
Racine	0	0		0	111	0	0	0	3	0	0	9
Superior	2	0		0	188	0	0	0	0	0	0	1
WEST NORTH CENTRAL												
Minnesota:					6.55							
Duluth	0	0		0	329	0	0 2	0	9	0	0	i
Minneapolis	0	0		0	37 67	1 0	4	0	9	0	0	8
Missouri:		-			-							
Kansas City	0	0	7	0	38	1	9	0	1	0	1	10
St. Joseph	0	0	1	0	320	0	13	0	8	0	0	12

[•] In some instances the figures include nonresident cases.

City reports for week ended Apr. 10, 1948-Continued

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	23868	tis, in-	Influ	enza	92	ens,	nia	itis	ver	868	and	ongh
Division, State, and city	Diphtheria cases	Encephalitis, fectious, case	Савев	Deaths	Measles cases	Meningitis, meningococcus, cases	P n e u m o r deaths	Poliomyelitis cases	Scarlet-fe cases	Smallpox cases	Typhoid-and paratyphoid fever cases	Whooping cough
WEST NORTH CENTRAL—												
Nebraska: Omaha Kansas:	0	0		0	95	0	4	0	1	0	0	
Topeka	0	0		0	15 6	0	1 4	0	0 2	0	0	
Deleware:												
Wilmington Maryland:	0	0		0	30	0	1	0	2	0	0	
Baltimore Cumberland Frederick	0 1	0	1	0 0	56	0 0	13 0 0	0 0	9	0	0 1 0	
District of Columbia: Washington	0	0		0	123	2	5	0	5	0	0	ı
Virginia: Lynchburg Richmond	0	0		0	1	0	6	0	0 3	0	0	3
Roanoke West Virginia: Charleston	0	0		0	3	0	1	0	1	0	0	·····i
Wheeling North Carolina: Raleigh	0	0		0	11	0	0	0	0	0	0	
Wilmington	0	0		0		0	0 2	0 2	0	0	0	3
Charleston	0	0	4	0	1	0	0	0	2	0	0	8
Atlanta Brunswick	1 0	0	2	1	2	0	2	0	2 0	0	0	1
Savannah Florida: Tampa	0	0		0	8	1	2 2	0	3	0	0	1
EAST SOUTH CENTRAL												
Tennessee: Memphis Nashville	0	0	4	1 0	166	0	8	0	8	0	0	1 3
Alabama: Birmingham	0	0	1	0	6	0	2	0	3	0	0	2
Mobile	0	0		0		0	1	0	0	0	0	
Arkansas: Little Rock	0	0	8	0	2	0	2	0	0	0	0	1
New Orleans	0	0	1	0		0	5	0	1	0	0	
Oklahoma: Oklahoma City Texas:	0	0		1	7	0	5	0	1	0	0	2
Dallas Galveston	0	1 0		0	131	0	0	0	6	0	0	1
HoustonSan Antonio	0	. 0	3	0 2	29	0	6	0	0	0	0 .	1
Montana:												
Billings Great Falls	0	0		0	3	0	1 2	0	4	0	0	
Helena Missoula daho:	0	0		0		0	0	0	0	0	0	
Boise	0	0		0		0	3	0	0	0	0 .	
Denver	0	0	1	0	309 16	0	5	0	6	0	0	17 5
Salt Lake City	0	0		1	19	0	3	0	3	0	0	1

City reports for week ended Apr. 10, 1948-Continued

	cases	s, in-	Influ	enza	2	me cus,	nia	litis	ever	cases	and	cough
Division, State, and city	Diphtheria	Encephalitis, fectious, cas	Cases	Deaths	Measles cases	Meningitis, me ningococcus, cases	Pneumo deaths	Poliomyeliti cases	Scarlet fo	Smallpox ca	Typhoid paratyph fever cases	00 00
PACIFIC .												
Washington:												
Seattle	0 0	0		0	38	0	3 2 0	0	6 2	0	0	4
Spokane	0	0		0	3		2	0	0	0	0	
TacomaCalifornia:	U	0		0	21	0	0	0	U	U	0	
Los Angeles	1	0	16	1	191	1	0	2	20	0	1	14
Sacramento	Ô	0		Ô	23	1 0	0 5	0	2	0	Ô	4
San Francisco	0	0	7	0	356	0	5	0	9	0	0	7
Total	36	1	64	15	7, 589	25	353	6	586	0	6	293
Corresponding week, 1947	63		796	69	1, 393		480		692	0	12	523
A verage 1943-471	69		208		37, 201		2 405		1, 576	ĭ	12	635

¹ Exclusive of Oklahoma City.

Rates (annual basis) per 100,000 population, by geographic groups, for the 89 cities in the preceding table (latest available estimated population, 34,593,800)

	case	in- case	Influ	ienza	rates	me-	death	0880	0830	rates	para- ever	cough
	Diphtheria rates	Encephalitis, fectious, rates	Case rates	Death rates	Measles case	Meningitis, ningococcus, rates	Pneumonia rates	Poliomyelitis rates	Scarlet fever	Smallpox case rates	Tyhpoid and typhoid f	Whooping conserates
New England	2.6	0.0	5.3	2.6	1, 331	0.0	89. 3	0.0	221	0.0	0.0	45
Middle Atlantic East North Central	6.0	0.0	3. 7 0. 6	1.4	1, 142 1, 299	3.7	54. 6 38. 9	0.5	77 116	0. 0	0.0	45 30 46 70 64 35
West North Central	10. 1	0.0	16.1	0.0	1, 826	4.0	74. 4	0.0	70	0.0	4.0	70
South Atlantic	8. 2	0.0	11.4	3.3	391	8.2	55.6	3.3	49	0.0	1.6	64
East South Central	0.0	0.0	29. 5	5.9	1,021	0.0	70.8	0.0	100	0.0	0.0	35
West South Central	7.6	2.5	22. 9	10. 2	452	5. 1	73.7	2.5	23	0.0	5.1	13
Mountain Pacific	7.9	0.0	7. 9 36. 4	7.9	2, 756 1, 000	0.0	119. 1 15. 8	0.0 3.2	127 62	0.0	0.0	183 46
Total	5. 4	0.2	9.7	2.3	1, 147	3.8	53. 4	0.9	89	0.0	0.9	44

Dysentery, amebic.—Cases: Memphis 1, New Orleans 1, Los Angeles 4, San Francisco 1.
Dysentery, bacillary.—Cases: Worcester 3, St. Louis 1, Los Angeles 2.
Dysentery, unspecified.—Cases: San Antonio 1.
Tularenia.—Cases: New Orleans 1, San Antonio 1.
Typhus fever, endemic.—Cases: New York 2, Tampa 1.

² 3-year average 1945-47. ³ 5-year median 1943-47.

TERRITORIES AND POSSESSIONS

Panama Canal Zone

Notifiable diseases-February 1948.-During the month of February 1948, certain notifiable diseases were reported in the Panama Canal Zone and terminal cities as follows:

	Residence ¹											
Disease	Panama City		Colon		Canal Zone		Outside the Zone and terminal cities		Total			
	Cases	Deaths	Cases	Deaths	Cases	Deaths	Cases	Deaths	Cases	Deaths		
Chickenpox	7 11				2		1		10 13			
Amebic Bacillary Malaria ²	i		2				148	1	1 8 157			
Measles	3	9		2	1 17	2	13	4	16 1 17	ii		
Poliomyelitis Relapsing fever Tuberculosis Typhoid fever	1	17		9	2	3		7	1 1 2	36		

If place of infection is known, cases are so listed instead of by residence.
 le recurrent cases.
 Reported in the Canal Zone only.

523 635

ties

case rates

DEATHS DURING WEEK ENDED APR. 10, 1948

[From the Weekly Mortality Index, issued by the National Office of Vital Statistics]

	Week ended Apr. 10, 1948	Correspond- ing week, 1947
Data for 93 large cities of the United States: Total deaths. Median for 3 prior years. Total deaths, first 15 weeks of year. Deaths under 1 year of age. Median for 3 prior years. Deaths under 1 year of age, first 15 weeks of year. Data from industrial insurance companies:	9, 690 9, 154 152, 347 714 599 10, 422	10, 154 151, 812 723 12, 075
Policies in force. Number of death claims. Death claims per 1,000 policies in force, annual rate. Death claims per 1,000 policies, first 15 weeks of year, annual rate	71, 084, 296 15, 613 11. 5 10. 3	67, 308, 805 12, 738 9, 9 9, 9

FOREIGN REPORTS

CANADA

Provinces—Communicable diseases—During the weeks ended March 20 and 27, 1948, cases of certain communicable diseases were reported by the Dominion Bureau of Statistics of Canada as follows:

Week ended March 20, 1948 .-

Disease	Prince Edward Island	Nova Scotia	New Bruns- wick	Que- bec	Onta- rio	Mani- toba	Sas- katch- ewan	Al- berta	British Colum- bia	Tota
Chickenpox		42		196	345	48	12	43	85	771
Diptheria				10				5	1	15
Dysentery, bacillary German measles				73	21	1	2	7	6	110
Influenza		41	1	881	20	1 2	7	15	19 146	2, 326
Measles Meningitis, meningococ-			1	881	1, 276	2		15	140	4, 34
cus			1	1	2	1				
Mumps Poliomyelitis		23		283	343	47	66	47	22	831
Scarlet fever			2	47	82	4		5	36	176
Tuberculosis (all forms)		3	4	87	37	30	9	3	36	206
Typhoid and paraty- phoid fever				15			1	1		17
Undulant fever				1	2		1	1	1	€
Venereal diseases: Gonorrhea	4	17	7	75	69	33	18	33	73	329
Syphilis	1	13	7 7	85	34	8	9	8	22	187
Other forms				25	26	2	1	55	5	114

Week ended March 27, 1948 .-

Disease	Prince Edward Island	Nova Scotia	New Bruns- wick	Que- bec	On- tario	Mani- toba	Sas- katch- ewan	Al- berta	British Colum- bia	Total
Chickenpox		34 1	3 1	171	264	60	20 1	13 1	89	654 12
German measles Influenza	*******	14		9	18 16			2	12 34	41 64
Measles Meningitis, meningococ- cus				797	1, 231	1	6	13	61	2, 112
Mumps		23	1	205	288	25	65	32	10	649
Scarlet fever		1	2	34	67	1	4	4	6	119
Tuberculosis (all forms) Typhoid and paraty-	*******	4	6	52	31	13	5	60	53	224
phoid fever Undulant fever				9		1			1 2	11
Venereal diseases:							***			332
		8	9	98	84	30	18	45	40	
7772	********		10	76 26	28 15	8 3	11	33	12 5	153 82

MADAGASCAR

Madagascar and Comoro Islands—Notifiable diseases—January and February 1948.—Notifiable contagious diseases were reported in Madagascar and Comoro Islands during January and February 1948 as follows:

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		Jan	uary		February					
	Aliens		Natives		Aliens		Natives			
	Cases	Deaths	Cases	Deaths	Cases	Deaths	Cases	Deaths		
Beri-beri	0	0	20	0	1	0	9	1		
Bilharziasis	1	0	86	0	0	0	114	(
Cerebrospinal meningitis	0	0	9	1	0	0	12	1 7		
Diphtheria	2	0	8	2	3	0	2	1		
Dysentery amebic	23	0	400	11	20	0	598	14		
Dysentery, bacillary	0	0	21	0	2	0	11	1		
Ervsipelas	1	0	10	1	0	0	18	1		
Influenza	4	0	2, 330	28	5	0	2, 247	2!		
Leprosy	0	0	52	0	0	0	25	(
Malaria	471	7	41, 644	374	564	4	41, 906	369		
Measles	2	0	57	2	1	0	39	(
Mumps	2	0	254	0	0	0	182	(
Plague	0	0	61	41	0	0	46	25		
Pneumonia, broncho	3	2	295	57	5	4	308	36		
Pneumonia, pneumococcic	5	1	419	79	2	1	293	56		
Poliomyelitis	0	0	0	0	0	0	2	(
Puerperal infection	0	0	4	1	0	0	8	0		
Relapsing fever	0	0	0	0	1	0	0			
Scarlet fever	1	0	Õ	0 1	1	0	0	1		
Trachoma	0	0	ĩ	0	î	0	0	1 6		
Tuberculosis, pulmonary	3	i	129	36	11	3	113	20		
Typhoid and paratyphoid fever	ĭ	0	48	12	3	1	30	6		
Whooping cough	6	0	82	0	2	0	88	0		

REPORTS OF CHOLERA, PLAGUE, SMALLPOX, TYPHUS FEVER, AND YELLOW FEVER RECEIVED DURING THE CURRENT WEEK

Note.—Except in cases of unusual incidence, only those places are included which had not previously reported any of the above-mentioned diseases, except yellow fever, during recent months. All reports of yellow fever are published currently.

A table showing the accumulated figures for these diseases for the year to date is published in the Public Health Reports for the last Friday in each month.

Cholera

India—Calcutta.—For the week ended April 3, 1948, 282 cases of cholera were reported in Calcutta, India.

Indochina (French)—Cochinchina—Rachgia.—During the period March 1-20, 1948, 58 cases of cholera with 23 deaths were reported in Rachgia, Cochinchina French Indo-China.

Plague

Belgian Congo—Stanleyville Province.—During the week ended April 3, 1948, 1 fatal case of plague was reported in the area northeast of Blukwa in Stanleyville Province, Belgian Congo.

British East Africa—Tanganyika—Central Province.—During the months of February and March 1948, 189 cases of plague with 85

deaths were reported in Singida District, Central Province, Tanganyika. British East Africa.

China—Kiangsi Province—Nanchang.—Information dated April 15, 1948, states that the usual spring epidemic of plague has re-appeared in Nanchang, Kiangsi Province, China. The epidemic is reported under control, but travelers have been advised to be innoculated.

India—Lucknow.—For the week ended March 27, 1948, 24 cases of plague with 8 deaths were reported in Lucknow, India.

Portugal—Azores.—For the period February 1-14, 1948, 3 cases of plague with 2 deaths were reported in the Ponta Delgada area, Azores, Portugal (4 cases with 1 death were reported in January 1948).

Smallpox

China—Shanghai.—For the week ended April 3, 1948, 94 cases of smallpox were reported in Shanghai, China.

France—Seine Department.—During the period February 16-March 31, 1948, 3 cases of smallpox were reported in Seine Department, France.

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India—Calcutta.—For the week ended April 3, 1948, 212 cases of smallpox were reported in Calcutta, India.

Portugal—Lisbon.—For the period March 1-27, 1948, 25 cases of smallpox with 2 deaths were reported in Lisbon, Portugal.

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It contains (1) current information regarding the incidence and geographic distribution of communicable diseases in the United States, insofar as data are obtainable, and of cholera, plague, smallpox, typhus fever, yellow fever, and other important communicable diseases throughout the world; (2) articles relating to the cause, prevention, and control of disease; (3) other pertinent information regarding sanitation and the conservation of the public health.

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